

SEARCH REQUEST FORM 52840 *msj*Examiner # (Mandatory): 77006 Requester's Full Name: JENNIFER HUNTArt Unit 1142 Location (Bldg/Room#): CN/8E10 (ex's mailbox) Phone (circle 305 306 308) 7548Serial Number: 09/186,475 Results Format Preferred (circle): PAPER DISK E-MAILTitle of Invention METHODS OF DETERMINING AN EFFICACIOUS DOSE OF A DRUGInventors (please provide full names): Annie Fong, Klaus HirthEarliest Priority Date: NOV. 7 1997

Keywords (include any known synonyms registry numbers, explanation of initialisms):

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## Search Topic:

Please write detailed statement of the search topic, and the concept of the invention. Describe as specifically as possible the subject matter to be searched. Define any terms that may have a special meaning. Give examples of relevant citations, authors, etc., if known. You may include a copy of the abstract and the broadcast or most relevant claim(s).

PLEASE SEARCH THE ATTACHED GENERIC COMPOUND (CLAIM 7) AND THE ATTACHED SPECIES (A). IN PARTICULAR THEY ARE CLAIMED AS ANGIOGENESIS INHIBITORS, OR ANGIOGENESIS RECEPTOR (FH-1 or FLK-1) ANTAGONISTS.

THX.  
*Jennifer*

## STAFF USE ONLY

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Searcher Location: \_\_\_\_\_

Date Picked Up: \_\_\_\_\_

Date Completed: 10/16/01

Clerical Prep Time: \_\_\_\_\_

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Number of Databases: \_\_\_\_\_

## Type of Search:

\_\_\_\_ N.A. Sequence

\_\_\_\_ A.A. Sequence

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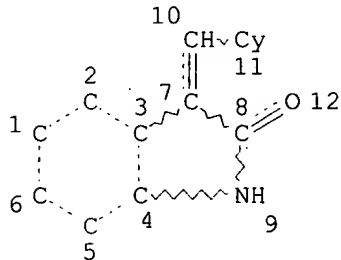
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 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:  
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 NUMBER OF NODES IS 12

STEREO ATTRIBUTES: NONE  
 L5 4211 SEA FILE=REGISTRY SSS FUL L3  
 L7 208 SEA FILE=HCAPLUS ABB=ON PLU=ON L5  
 L11 361 SEA FILE=REGISTRY ABB=ON PLU=ON ANGIOGEN? OR FH1 OR FLK?  
 L12 17556 SEA FILE=HCAPLUS ABB=ON PLU=ON L11 OR ?ANGIOGEN? OR FH1 OR  
 FLK? OR FH(W)1  
 L14 16 SEA FILE=HCAPLUS ABB=ON PLU=ON L7(L)L12

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L14 ANSWER 1 OF 16 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2001:416904 HCAPLUS

DOCUMENT NUMBER: 135:19497

TITLE: Preparation of curcumin analogues for treating cancer

INVENTOR(S): Snyder, James P.; Davis, Matthew C.; Adams, Brian;  
Shoji, Mamoru; Liotta, Dennis C.; Ferstl, Eva M.;  
Sunay, Ustun B.

PATENT ASSIGNEE(S): Emory University, USA

SOURCE: PCT Int. Appl., 69 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

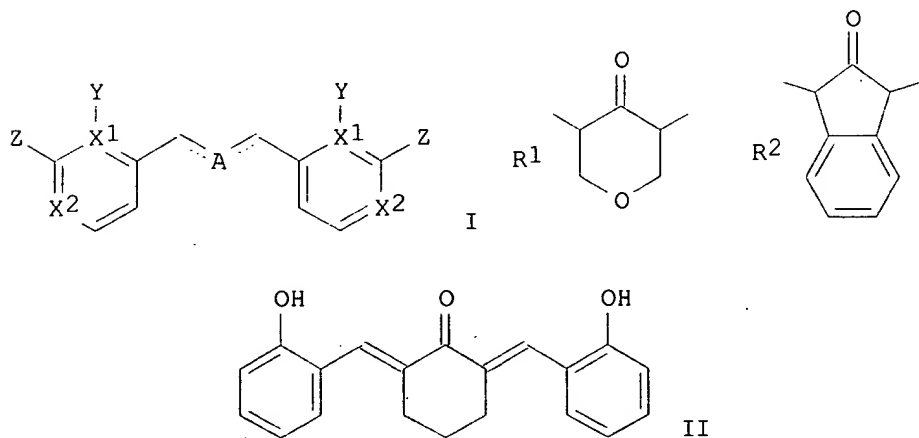
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001040188	A1	20010607	WO 2000-US32870	20001204
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
PRIORITY APPLN. INFO.:			US 1999-168913	P 19991203
OTHER SOURCE(S):			MARPAT 135:19497	

GI



AB Curcumin analogs of formula I [Y = OH, halo, CF<sub>3</sub>; Z = H, (substituted) OH, halo, CF<sub>3</sub>; X1, X2 = C, N; A = CHCOCH, NHCONH, CHCH(OH)CH, R1, R2, etc.]

are prepd. which exhibit antitumor and anti-angiogenic properties. Thus, II was prepd. from cyclohexanone and 2-hydroxybenzaldehyde. The growth inhibitory concn. of II in the NCI anti-tumor screen was lower than CISPLATIN for several cell types.

IT 3367-88-2P 342808-37-1P 342808-38-2P

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of curcumin analogs as antitumor and anti-angiogenic agents)

REFERENCE COUNT: 2

REFERENCE(S): (1) El-Subbagh; J MED CHEM 2000, V43(15), P2915 HCAPLUS  
(2) Teuscher, E; PHARMAZIE 1987, V42(2), P109 HCAPLUS

L14 ANSWER 2 OF 16 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2001:169669 HCAPLUS

DOCUMENT NUMBER: 134:348057

TITLE: The antiangiogenic protein kinase inhibitors SU5416 and SU6668 inhibit the SCF receptor (c-kit) in a human myeloid leukemia cell line and in acute myeloid leukemia blasts

AUTHOR(S): Smolich, Beverly D.; Yuen, Helene A.; West, Kristina A.; Giles, Francis J.; Albitar, Maher; Cherrington, Julie M.

CORPORATE SOURCE: Sugan, South San Francisco, CA, 94080, USA

SOURCE: Blood (2001), 97(5), 1413-1421  
CODEN: BLOOAW; ISSN: 0006-4971

PUBLISHER: American Society of Hematology

DOCUMENT TYPE: Journal

LANGUAGE: English

AB SU5416 and SU6668 are potent antiangiogenic small-mol. inhibitors of receptor tyrosine kinases, including those of the vascular endothelial growth factor and platelet-derived growth factor receptor families. The stem cell factor (SCF) receptor, c-kit, is structurally related to these receptors and, although not expressed on mature peripheral blood cells, is expressed in leukemic blasts derived from 60% to 80% of acute myeloid leukemia (AML) patients. The c-kit kinase inhibitory activity of SU5416 and SU6668 was evaluated in MO7E cells, a human myeloid leukemia cell line. Tyrosine autophosphorylation of the receptor, induced by SCF, was inhibited in these cells by SU5416 and SU6668 in a dose-dependent manner (inhibitory concn. of 50% [IC50] 0.1-1 .mu.M). Inhibition of extracellular signal-regulated kinase 1/2 (ERK1/2) phosphorylation, a signaling event downstream of c-kit activation, was also inhibited in a dose-dependent manner. Both compds. also inhibited SCF-induced proliferation of MO7E cells (IC50 0.1 .mu.M for SU5416; 0.29 .mu.M for SU6668). Furthermore, both SU5416 and SU6668 induced apoptosis in a dose- and time-dependent manner as measured by the increase in activated caspase-3 and the enhanced cleavage of its substrate poly(ADP-ribose) polymerase. These findings with MO7E cells were extended to leukemic blasts from c-kit+ patients. In patient blasts, both SU5416 and SU6668 inhibited SCF-induced phosphorylation of c-kit and ERK1/2 and induced apoptosis. These studies indicate that SU5416 and SU6668 inhibit biol. functions of c-kit in addn. to exhibiting antiangiogenic properties and suggest that the combination of these activities may provide a novel therapeutic approach for the treatment of AML.

IT 204005-46-9, SU5416

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(the **antiangiogenic** protein kinase inhibitors SU5416 and SU6668 inhibit the SCF receptor (c-kit) in a human myeloid leukemia cell line and in acute myeloid leukemia blasts)

REFERENCE COUNT: 27  
 REFERENCE(S): (1) Aguayo, A; Blood 1999, V94, P3717 HCAPLUS  
 (3) Bellamy, W; Cancer Res 1999, V59, P728 HCAPLUS  
 (4) Bene, M; Blood 1998, V92, P596 HCAPLUS  
 (6) Chabot, B; Nature 1988, V335, P88 HCAPLUS  
 (7) Dias, S; J Clin Invest 2000, V106, P511 HCAPLUS  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 3 OF 16 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2001:96419 HCAPLUS  
 DOCUMENT NUMBER: 135:116407  
 TITLE: Zebrafish angiogenesis: A new model for drug screening  
 AUTHOR(S): Serbedzija, George N.; Flynn, Edward; Willett, Catherine E.  
 CORPORATE SOURCE: Phylionix Pharmaceuticals, Inc., Cambridge, MA, 02139, USA  
 SOURCE: Angiogenesis (2000), Volume Date 1999, 3(4), 353-359  
 CODEN: AGIOFT; ISSN: 0969-6970  
 PUBLISHER: Kluwer Academic Publishers  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English

AB Angiogenesis is necessary for tumor growth, making inhibition of vessel formation an excellent target for cancer therapy. Current assays for angiogenesis, however, are too complex to be practical for drug screening. Here, we demonstrate that the zebrafish is a viable whole animal model for screening small mols. that affect blood vessel formation. Blood vessel patterning is highly characteristic in the developing zebrafish embryo and the subintestinal vessels (SIVs) can be stained and visualized microscopically as a primary screen for compds. that affect angiogenesis. Small mols. added directly to the fish culture media diffuse into the embryo and induce observable, dose-dependent effects. To evaluate the zebrafish as a model, we used two angiogenesis inhibitors, SU5416 and TNP470, both of which have been tested in mammalian systems. Both compds. caused a redn. in vessel formation when introduced to zebrafish embryos prior to the onset of angiogenesis. Short duration (1 h) exposure of SU5416 was sufficient to block new angiogenic and vasculogenic vessel formation. In contrast, TNP470 required continuous exposure to block SIV formation and had no apparent effect on vasculogenic vessel formation. To ascertain whether blood vessels in the zebrafish embryo respond to angiogenic compds., we introduced human VEGF into embryos. Injection of VEGF caused an observable increase in SIV formation.

IT 204005-46-9, SU5416

RL: BAC (Biological activity or effector, except adverse); BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)  
 (zebrafish **angiogenesis**: a model for drug screening)

REFERENCE COUNT: 22  
 REFERENCE(S): (2) Bergers, G; Science 1999, V284, P808 HCAPLUS  
 (3) Detrich, H; Proc Natl Acad Sci USA 1995, V92, P10713 HCAPLUS  
 (4) Drake, C; Proc Natl Acad Sci USA 1995, V92, P7657 HCAPLUS  
 (5) D'Amore, P; Angiogenesis Sci Med 1999, P44 HCAPLUS  
 (6) Ferrara, N; Endocr Rev 1997, V18, P4 HCAPLUS  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 4 OF 16 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2001:83373 HCAPLUS  
 DOCUMENT NUMBER: 135:86674  
 TITLE: The angiogenesis inhibitor SU5416 has long-lasting effects on vascular endothelial growth factor receptor phosphorylation and function  
 AUTHOR(S): Mendel, Dirk B.; Schreck, Randall E.; West, David C.; Li, Guangmin; Strawn, Laurie M.; Tanciongco, Sheila S.; Vasile, Stefan; Shawver, Laura K.; Cherrington, Julie M.  
 CORPORATE SOURCE: Sugan, Inc., South San Francisco, CA, 94080, USA  
 SOURCE: Clin. Cancer Res. (2000), 6(12), 4848-4858  
 CODEN: CCREF4; ISSN: 1078-0432  
 PUBLISHER: American Association for Cancer Research  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English

AB SU5416, a selective inhibitor of the tyrosine kinase activity of the vascular endothelial growth factor (VEGF) receptor Flk-1/KDR, is currently in Phase III clin. trials for the treatment of advanced malignancies. In cellular assays, SU5416 inhibits the VEGF-dependent mitogenic/proliferative response of human umbilical vein endothelial cells (HUVECs). In tumor xenograft models, SU5416 inhibits the growth of tumors from a variety of origins by inhibiting tumor angiogenesis. In three different human tumor xenograft models, infrequent (once or twice a week) administration of SU5416 is efficacious despite the fact that it has a short plasma half-life (30 min), which suggests that SU5416 has long-lasting inhibitory activity in vivo. The goal of the present study was to det. the basis for the prolonged activity of SU5416. The results indicate that a short (3 h) exposure to 5 .mu.M SU5416 (to mimic plasma levels of the compd. as measured in patients who were receiving SU5416 therapy) produced long-lasting (at least 72 h) inhibition of the VEGF-dependent proliferation of HUVECs in culture, which indicate that SU5416 has long-lasting inhibitory activity in vitro as well as in vivo. SU5416 treatment of HUVECs did not affect surface expression of Flk-1/KDR or the affinity of the receptor for VEGF. Instead, the durability of the in vitro activity of SU5416 was shown to be attributable to its long-lasting ability to specifically inhibit VEGF-dependent phosphorylation of Flk-1/KDR and subsequent downstream signaling, although SU5416 is not an irreversible inhibitor of Flk-1/KDR tyrosine kinase activity. The long-lasting inhibition of cellular responses to VEGF was attributable to the accumulation of SU5416 in cells, as shown using radiolabeled compd., such that inhibitory cellular concns. of SU5416 are maintained long after the removal of the compd. from the medium. The long-lasting inhibitory activity of SU5416 in vitro is consistent with the finding that SU5416 has demonstrated evidence of biol. activity in clin. studies when administered twice a week despite a short plasma half-life.

IT 204005-46-9, SU5416

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (angiogenesis inhibitor SU5416 has long-lasting effects on vascular endothelial growth factor receptor phosphorylation and function in relation to antitumor effects)

REFERENCE COUNT: 48

REFERENCE(S): (2) Asano, M; Cancer Res 1995, V55, P5296 HCAPLUS  
 (5) Bridges, A; Curr Med Chem 1999, V6, P825 HCAPLUS  
 (6) Cherrington, J; Adv Cancer Res 2000, V79, P1 HCAPLUS  
 (7) Claffey, K; Cancer Res 1996, V56, P172 HCAPLUS  
 (9) Discafani, C; Biochem Pharmacol 1999, V57, P917 HCAPLUS

## ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 5 OF 16 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:860816 HCAPLUS

DOCUMENT NUMBER: 134:141334

TITLE: Biotransformation of the anti-angiogenic compound  
SU5416AUTHOR(S): Antonian, Lida; Zhang, Hongbing; Yang, Cheng; Wagner,  
Greg; Shawver, Laura K.; Shet, Manjunath; Ogilvie,  
Brian; Madan, Ajay; Parkinson, Andrew

CORPORATE SOURCE: SUGEN, Inc., South San Francisco, CA, 94080, USA

SOURCE: Drug Metab. Dispos. (2000), 28(12), 1505-1512

CODEN: DMDSAI; ISSN: 0090-9556

PUBLISHER: American Society for Pharmacology and Experimental  
Therapeutics

DOCUMENT TYPE: Journal

LANGUAGE: English

AB SU5416 [3-(3,5-dimethyl-1H-pyrrol-2-ylmethylene)-1,3-dihydro-indol-2-one], an inhibitor of VEGF (vascular endothelial growth factor) receptor tyrosine kinase, Flk-1/KDR (fetal liver kinase 1/kinase insert domain-contg. receptor), also known as VEGF receptor 2 (VEGFR2) is in advanced clin. trials for treatment of AIDS-related Kaposi's sarcoma and colorectal and nonsmall cell lung cancers. Since this chem. class has not been studied previously with therapeutic intent, the present study was designed to investigate the in vitro metab. of SU5416 by mouse, rat, dog, monkey, and human liver microsomes and to identify the major metabolites of SU5416. An HPLC procedure was developed and validated to resolve and quantify SU5416 and its metabolites. To evaluate the in vitro metab. of SU5416, pooled liver microsomes from mice, rats, dogs, monkeys, and humans were incubated with SU5416 (25 .mu.M) in the presence of an NADPH-generating system. In the presence of NADPH, mouse, rat, dog, monkey, and human liver microsomes converted SU5416 to at least 12, 9, 9, 7, and 6 polar metabolites, resp. Microsomal metab. of SU5416 showed marked species differences in the levels of different metabolites formed. The overall rate of SU5416 metab. by liver microsomes from the species examd. followed the rank order: monkey .gtoreq. mouse .apprxeq. rat > dog > human. Two major metabolites of SU5416 were identified, a hydroxymethyl deriv. of SU5416 (M12) and a carboxylic acid deriv. of SU5416 (M6), by spectroscopic methods and comparison with authentic compds. Both of these oxidative metabolites were further metabolized in vivo through glucuronidation. The metabolic fate of SU5416 in microsomes from various species as well as data from in vivo biotransformation in the rat are discussed.

IT 204005-46-9, SU5416

RL: ANT (Analyte); BPR (Biological process); ANST (Analytical study); BIOL (Biological study); PROC (Process)

(biotransformation of anti-angiogenic compd. SU5416)

IT 280748-38-1, SU 6689 280748-39-2, SU 6595

280748-41-6, SU 9838 324047-04-3 324047-05-4

RL: MFM (Metabolic formation); BIOL (Biological study); FORM (Formation, nonpreparative)

(biotransformation of anti-angiogenic compd. SU5416)

REFERENCE COUNT: 19

REFERENCE(S): (1) Abulafia, O; Gynecol Oncol 1999, V72, P220 HCAPLUS  
(2) Cherrington, J; Adv Cancer Res 2000, V79, P1 HCAPLUS  
(5) Ferrara, N; J Mol Med 1999, V77, P527 HCAPLUS  
(6) Folkman, J; Nat Med 1995, V1, P27 HCAPLUS  
(7) Fong, T; Cancer Res 1999, V59, P99 HCAPLUS

## ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 6 OF 16 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:685884 HCAPLUS

DOCUMENT NUMBER: 133:308083

TITLE: Inhibition of angiogenesis decreases alveolarization in the developing rat lung

AUTHOR(S): Jakkula, Malathi; Le Cras, Timothy D.; Gebb, Sarah; Hirth, K. Peter; Tudor, Rubin M.; Voelkel, Norbert F.; Abman, Steven H.

CORPORATE SOURCE: Pediatric Pulmonary Medicine, Pediatric Heart Lung Center, Department of Pediatrics, University of Colorado School of Medicine, Denver, CO, 80218, USA

SOURCE: Am. J. Physiol. (2000), 279(3, Pt. 1), L600-L607

CODEN: AJPHAP; ISSN: 0002-9513

PUBLISHER: American Physiological Society

DOCUMENT TYPE: Journal

LANGUAGE: English

AB To det. whether angiogenesis is necessary for normal alveolarization, the authors studied the effects of two antiangiogenic agents, thalidomide and fumagillin, on alveolarization during a crit. period of lung growth in infant rats. Newborn rats were treated with daily injections of fumagillin, thalidomide, or vehicle during the first 2 wk of life. Compared with control treatment, fumagillin and thalidomide treatment reduced lung wt.-to-body wt. ratio and pulmonary arterial d. by 20 and 36%, resp., and reduced alveolarization by 22%. Because these drugs potentially have nonspecific effects on lung growth, the authors also studied the effects of Su-5416, an inhibitor of the vascular endothelial growth factor receptor known as kinase insert domain-contg. receptor/fetal liver kinase (KDR/flk)-1. As obsd. with the other antiangiogenic agents, Su-5416 treatment decreased alveolarization and arterial d. The authors conclude that treatment with three different antiangiogenic agents attenuated lung vascular growth and reduced alveolarization in the infant rat. The authors speculate that angiogenesis is necessary for alveolarization during normal lung development and that injury to the developing pulmonary circulation during a crit. period of lung growth can contribute to lung hypoplasia.

IT 204005-46-9, Su-5416

RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)  
(**antiangiogenic** treatment decreases alveolarization in developing rat lung)

REFERENCE COUNT: 51

REFERENCE(S): (2) Acarregui, M; Am J Respir Cell Mol Biol 1999, V20, P14 HCAPLUS  
(3) Babaei, S; Circ Res 1998, V82, P1007 HCAPLUS  
(4) Beck, L; FASEB J 1997, V11, P365 HCAPLUS  
(5) Blanco, L; Am J Physiol Lung Cell Mol Physiol 1989, V257, PL240 HCAPLUS  
(12) DeMello, D; Am J Respir Cell Mol Biol 1997, V16, P568 HCAPLUS

## ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 7 OF 16 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:509842 HCAPLUS

DOCUMENT NUMBER: 133:348326

TITLE: Targeting angiogenesis inhibits tumor infiltration and expression of the pro-invasive protein SPARC

AUTHOR(S): Vajkoczy, Peter; Menger, Michael D.; Goldbrunner, Roland; Ge, Shugang; Annie, T.; Fong, T.; Vollmar,



Brigitte; Schilling, Lothar; Ullrich, Axel; Hirth, K. Peter; Tonn, Jorg C.; Schmiedek, Peter; Rempel, Sandra A.

## CORPORATE SOURCE:

Department of Neurosurgery. Klinikum Mannheim, University of Heidelberg, Mannheim, D-68167, Germany

## SOURCE:

Int. J. Cancer (2000), 87(2), 261-268

CODEN: IJCNAW; ISSN: 0020-7136

## PUBLISHER:

Wiley-Liss, Inc.

## DOCUMENT TYPE:

Journal

## LANGUAGE:

English

AB The solid growth of high-grade glioma appears to be critically dependent on tumor angiogenesis. It remains unknown, however, whether the diffuse infiltration of glioma cells into healthy adjacent tissue is also dependent on the formation of new tumor vessels. Here, the authors analyze the relation between tumor angiogenesis and tumor cell infiltration in an exptl. glioma model. C6 cells were implanted into the dorsal skinfold chamber of nude mice, and tumor angiogenesis was monitored by intravital fluorescence videomicroscopy. Glioma infiltration was assessed by the extent of tumor cell invasion into the adjacent chamber tissue and by expression of SPARC, a cellular marker of glioma invasiveness. To test the hypothesis that glioma angiogenesis and glioma infiltration are codependent, the authors assessed tumor infiltration in both the presence and the absence of the angiogenesis inhibitor SU5416. SU5416 is a selective inhibitor of the VEGF/Flk-I signal-transduction pathway, a crit. pathway implicated in angiogenesis. Control tumors demonstrated both high angiogenic activity and tumor cell invasion accompanied by strong expression of SPARC in invading tumor cells at the tumor-host tissue border. SU5416-treated tumors demonstrated reduced vascular d. and vascular surface in the tumor periphery accompanied by marked inhibition of glioma invasion and decreased SPARC expression. A direct effect of SU5416 on glioma cell motility and invasiveness was excluded by in vitro migration and invasion assays. These results suggest a crucial role for glioma-induced angiogenesis as a prerequisite for diffuse tumor invasion and a possible therapeutic role for anti-angiogenic compds. as inhibitors of both solid and diffuse infiltrative tumor growth.

IT 204005-46-9, SU5416

RL: BAC (Biological activity or effector, except adverse); THU

(Therapeutic use); BIOL (Biological study); USES (Uses)

(targeting **angiogenesis** inhibits tumor infiltration and expression of pro-invasive protein SPARC using VEGF/Flk-I signal-transduction pathway inhibitor SU5416)

REFERENCE COUNT: 21

## REFERENCE(S):

- (2) Brooks, P; Cell 1994, V79, P1157 HCAPLUS
- (4) Fong, T; Cancer Res 1999, V59, P99 HCAPLUS
- (6) Kupprion, C; J Biol Chem 1998, V273, P29635 HCAPLUS
- (9) Millauer, B; Nature (Lond) 1994, V367, P576 HCAPLUS
- (10) Murphy-Ullrich, J; J Cell Biochem 1995, V57, P341 HCAPLUS

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 8 OF 16 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:459469 HCAPLUS

DOCUMENT NUMBER: 133:144316

## TITLE:

Development of SU5416, a selective small molecule inhibitor of VEGF receptor tyrosine kinase activity, as an anti-angiogenesis agent

## AUTHOR(S):

Mendel, Dirk B.; Laird, A. Douglas; Smolich, Beverly

D.; Blake, Robert A.; Liang, Congxin; Hannah, Alison L.; Shaheen, Raymond M.; Ellis, Lee M.; Weitman, Steve; Shawver, Laura K.; Cherrington, Julie M.  
 CORPORATE SOURCE: SUGEN, Inc., South San Francisco, CA, 94080, USA  
 SOURCE: Anti-Cancer Drug Des. (2000), 15(1), 29-41  
 CODEN: ACDDEA; ISSN: 0266-9536  
 PUBLISHER: Oxford University Press  
 DOCUMENT TYPE: Journal; General Review  
 LANGUAGE: English

AB A review with over 60 refs. Angiogenesis, or the sprouting of new blood vessels, is a central process in the growth of solid tumors. For many cancers, the extent of vascularization of a tumor is a neg. prognostic indicator signifying aggressive disease and increased potential for metastasis. Recent efforts to understand the mol. basis of tumor-assocd. angiogenesis have identified several potential therapeutic targets, including the receptor tyrosine kinases for the angiogenic factor vascular endothelial growth factor (VEGF). Here we review the approach taken at SUGEN, Inc. to discover and develop small mol. inhibitors of receptor tyrosine kinases as anti-angiogenic agents. We focus on SU5416, a selective inhibitor of VEGF receptors that is currently in clin. development for the treatment of advanced malignancies. Its biochem., biol. and pharmacol. properties are reviewed and clin. implications discussed.

IT 204005-46-9, SU5416  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (VEGF receptor tyrosine kinase inhibitors SU5416 development as anti-angiogenesis agent)

REFERENCE COUNT: 60  
 REFERENCE(S): (1) Adnane, J; Oncogene 1991, V6, P659 HCAPLUS  
 (2) Albin, A; Nature Medicine 1996, V2, P1371 HCAPLUS  
 (3) Angelov, L; Cancer Research 1999, V59, P5536 HCAPLUS  
 (5) Armesilla, A; Molecular and Cellular Biology 1999, V19, P2032 HCAPLUS  
 (6) Bais, C; Nature 1998, V391, P86 HCAPLUS  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 9 OF 16 HCAPLUS COPYRIGHT 2001 ACS  
 ACCESSION NUMBER: 2000:417312 HCAPLUS  
 DOCUMENT NUMBER: 133:159618  
 TITLE: Identification of Substituted 3-[(4,5,6,7-Tetrahydro-1H-indol-2-yl)methylene]-1,3-dihydroindol-2-ones as Growth Factor Receptor Inhibitors for VEGF-R2 (Flk-1/KDR), FGF-R1, and PDGF-R.beta. Tyrosine Kinases  
 AUTHOR(S): Sun, Li; Tran, Ngoc; Liang, Congxing; Hubbard, Steve; Tang, Flora; Lipson, Kenneth; Schreck, Randall; Zhou, Yong; McMahon, Gerald; Tang, Cho  
 CORPORATE SOURCE: SUGEN Inc., South San Francisco, CA, 94080-4811, USA  
 SOURCE: J. Med. Chem. (2000), 43(14), 2655-2663  
 CODEN: JMCMAR; ISSN: 0022-2623  
 PUBLISHER: American Chemical Society  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English

AB A series of new 3-substituted indolin-2-ones contg. a tetrahydroindole moiety was developed as specific inhibitors of receptor tyrosine kinases assocd. with VEGF-R, FGF-R, and PDGF-R growth factor receptors. These compds. were evaluated for their inhibitory properties toward VEGF-R2 (Flk-1/KDR), FGF-R1, PDGF-R.beta., p60c-Src, and EGF-R tyrosine kinases and their ability to inhibit growth factor-dependent cell proliferation.

Structure-activity relationships of this new pharmacophore have been detd. at the level of kinase inhibition. Compds. contg. a propionic acid moiety at the C-3' position of the tetrahydroindole ring represented the most potent indolin-2-ones to inactivate the VEGF, FGF, and PDGF receptor kinases. The inhibitory activities of 3-[3-(2-carboxyethyl)-4,5,6,7-tetrahydro-1H-indol-2-ylmethylene]-2-oxo-2,3-dihydro-1H-indole-5-carboxylic acid against VEGF-R2 (Flk-1), 3-[2-[6-(2-methoxyphenyl)-2-oxo-1,2-dihydroindol-3-ylidenemethyl]-4,5,6,7-tetrahydro-1H-indol-3-yl]propionic acid against FGF-R1, and 3-[2-(5-bromo-2-oxo-1,2-dihydroindol-3-ylidenemethyl)-4,5,6,7-tetrahydro-1H-indol-3-yl]propionic acid (I) against PDGF-R.beta. were 4, 80, and 4 nM, resp. However, all of these compds. were inactive when tested against the EGF-R tyrosine kinase. Compds. 3-[2-(2-oxo-1,2-dihydroindol-3-ylidenemethyl)-4,5,6,7-tetrahydro-1H-indol-3-yl]propionic acid (II) and I represented the most potent inhibitors of these classes to inhibit both biochem. kinase and growth factor-dependent cell proliferation for these three targets. In addn., compd. II was cocrystd. with the catalytic domain of FGF-R1 providing evidence to explain the structure-activity relation results. This study has provided evidence to support the potential of these new tyrosine kinase inhibitors for the treatment of angiogenesis and other growth factor-related diseases including human cancers.

IT 204005-46-9, Su5416 215543-92-3

RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study)

(substituted [(tetrahydroindolyl)methylene]dihydroindolones as growth factor receptor inhibitors for VEGF-R2 (Flk-1/KDR) and FGF-R1, and PDGF-R.beta. tyrosine kinases and as inhibitors of growth factor-dependent cell proliferation)

IT 288144-28-5P

RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); PRP (Properties); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); PROC (Process)

(substituted [(tetrahydroindolyl)methylene]dihydroindolones as growth factor receptor inhibitors for VEGF-R2 (Flk-1/KDR) and FGF-R1, and PDGF-R.beta. tyrosine kinases and as inhibitors of growth factor-dependent cell proliferation)

IT 288144-19-4P 288144-20-7P 288144-21-8P

288144-22-9P 288144-23-0P 288144-24-1P

288144-25-2P 288144-26-3P 288144-27-4P

288144-29-6P 288144-30-9P 288144-31-0P

288144-32-1P 288144-33-2P 288144-34-3P

288144-35-4P 288144-36-5P

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(substituted [(tetrahydroindolyl)methylene]dihydroindolones as growth factor receptor inhibitors for VEGF-R2 (Flk-1/KDR) and FGF-R1, and PDGF-R.beta. tyrosine kinases and as inhibitors of growth factor-dependent cell proliferation)

REFERENCE COUNT: 17

REFERENCE(S): (3) Folkman, J; J Biol Chem 1992, V267, P10931 HCAPLUS  
(6) Hanahan, D; Cell 1996, V86, P353 HCAPLUS  
(10) Mohammadi, M; Cell 1996, V86, P577 HCAPLUS  
(11) Mohammadi, M; Science 1997, V276, P955 HCAPLUS  
(12) Otwinowski, Z; Methods Enzymol 1997, V276, P307 HCAPLUS

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 10 OF 16 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:144722 HCAPLUS

DOCUMENT NUMBER: 132:185454  
 TITLE: Use of anti-angiogenic agents for inhibiting vessel wall injury  
 INVENTOR(S): Brown, Charles L., III; Gorlin, Steve  
 PATENT ASSIGNEE(S): Global Vascular Concepts, Inc., USA  
 SOURCE: PCT Int. Appl., 29 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000010552	A2	20000302	WO 1999-US19218	19990824
WO 2000010552	A3	20001123		

W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

AU 9956871	A1	20000314	AU 1999-56871	19990824
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PRIORITY APPLN. INFO.: US 1998-97579 P 19980824  
 WO 1999-US19218 W 19990824

AB Use of anti-angiogenic agents to inhibit an undesirable response to vessel wall injury, including stent neointima, dialysis graft neointima, vascular graft-induced neointima, and the treatment of benign hypertrophic scar formation as well as the treatment and passivation of unstable atherosclerotic plaques are provided. The invention provides for the use of catheter-based devices for enhancing the local delivery of anti-angiogenic agents into the endothelial tissues of blood vessels of the living body.

IT 204005-46-9, SU5416

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (anti-angiogenic agents for inhibiting vessel wall injury)

L14 ANSWER 11 OF 16 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:117816 HCAPLUS

DOCUMENT NUMBER: 133:37825

TITLE: Inhibition of tumor growth, angiogenesis, and microcirculation by the novel Flk-1 inhibitor SU5416 as assessed by intravital multifuorescence videomicroscopy

AUTHOR(S): Vajkoczy, Peter; Menger, Michael D.; Vollmar, Brigitte; Schilling, Lothar; Schmiedek, Peter; Hirth, K. Peter; Ullrich, Axel; Fong, T. Annie T.

CORPORATE SOURCE: Department of Neurosurgery, Klinikum Mannheim, University of Heidelberg, Mannheim, D-68167, Germany

SOURCE: Neoplasia (N. Y.) (1999), 1(1), 31-41

CODEN: NEOPFL; ISSN: 1522-8002

PUBLISHER: Stockton Press

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Vascular endothelial growth factor (VEGF) plays a fundamental role in mediating tumor angiogenesis and tumor growth. The direct effect of

SU5416, a novel small-mol. inhibitor of the Flk-1-mediated signal transduction pathway of VEGF, on tumor angiogenesis and microhemodynamics of an exptl. glioblastoma was investigated by intravital multifuorescence videomicroscopy. SU5416 treatment suppressed tumor growth. In parallel, SU5416 demonstrated a potent antiangiogenic activity, resulting in redn. of both the total and functional vascular d. of the tumor microvasculature, which indicates an impaired vascularization as well as perfusion failure in the treated tumors. This malperfusion was not compensated for by changes in vessel diam. or recruitment of nonperfused vessels. Analyses of the tumor microcirculation revealed microhemodynamic changes after angiogenesis blockade, such as a higher red cell velocity and blood flow in remnant tumor vessels than in controls. The results demonstrate that the novel antiangiogenic concept of targeting the tyrosine kinase of Flk-1/KDR by means of a small-mol. inhibitor represents an efficient strategy for controlling growth and progression of angiogenesis-dependent tumors.

IT 204005-46-9, SU 5416

RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study)

(inhibition of glioblastoma growth, **angiogenesis**, and microcirculation by tyrosine kinase inhibitor SU5416)

REFERENCE COUNT: 25

- REFERENCE(S):
- (1) Abramovitch, R; Cancer Res 1995, V55, P1956 HCAPLUS
  - (2) Andrade, S; Br J Pharmacol 1992, V107, P1092 HCAPLUS
  - (5) Boucher, Y; Cancer Res 1996, V56, P4264 HCAPLUS
  - (6) Brogi, E; J Clin Invest 1996, V97, P469 HCAPLUS
  - (7) Cheng, S; Proc Natl Acad Sci USA 1996, V93, P8502 HCAPLUS

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 12 OF 16 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1999:725101 HCAPLUS

DOCUMENT NUMBER: 132:30444

TITLE: Inhibition of angiogenesis by blocking activation of the vascular endothelial growth factor receptor 2 leads to decreased growth of neurogenic sarcomas

AUTHOR(S): Angelov, Lilyana; Salhia, Bodour; Roncari, Luba; McMahon, Gerald; Guha, Abhijit

CORPORATE SOURCE: Division of Neurosurgery, Toronto Western Hospital, University of Toronto, Toronto, ON, M5T 2S8, Can.

SOURCE: Cancer Res. (1999), 59(21), 5536-5541

CODEN: CNREA8; ISSN: 0008-5472

PUBLISHER: AACR Subscription Office

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Neurogenic sarcomas are incurable, common malignant human peripheral nerve tumors subject to local recurrence and systemic metastasis. In this study, the vascularity, vascular endothelial growth factor (VEGF) expression, and effects of inhibiting VEGF receptor on growth of neurogenic sarcomas were examd. Vascularization and VEGF expression were 6.4- and 15-fold higher in tumors than in normal nerves. The small mol. inhibitor (SU5416) of VEGF receptor 2 had no effect on neurogenic sarcoma cell lines in vitro, but the growth of a human tumor explant xenograft model was reduced by 54.8% compared to vehicle. Redn. in tumor growth was due to decreased tumor angiogenesis, leading to redn. of tumor cell proliferation and increased apoptosis. Inhibiting VEGF function may therefore be a useful adjuvant therapy for neurogenic sarcomas.

IT 204005-46-9, SU5416

RL: BAC (Biological activity or effector, except adverse); THU  
(Therapeutic use); BIOL (Biological study); USES (Uses)  
(angiogenesis inhibition by VEGF2 blocking leads to decreased  
growth of neurogenic sarcomas)

REFERENCE COUNT: 69

REFERENCE(S): (1) Adamis, A; Biochem Biophys Res Commun 1993, V193,  
P631 HCAPLUS  
(2) Alon, T; Nat Med 1995, V1, P1024 HCAPLUS  
(5) Basu, T; Nature 1992, V356, P713 HCAPLUS  
(6) Brown, L; Cancer Res 1993, V53, P4727 HCAPLUS  
(7) Cawthon, R; Cell 1990, V62, P193 HCAPLUS  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 13 OF 16 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1999:725076 HCAPLUS

DOCUMENT NUMBER: 132:44604

TITLE: Antiangiogenic therapy targeting the tyrosine kinase  
receptor for vascular endothelial growth factor  
receptor inhibits the growth of colon cancer liver  
metastasis and induces tumor and endothelial cell  
apoptosis

AUTHOR(S): Shaheen, Raymond M.; Davis, Darren W.; Liu, Wenbiao;  
Zebrowski, Brian K.; Wilson, Michael R.; Bucana,  
Corazon D.; McConkey, David J.; McMahon, Gerald;  
Ellis, Lee M.

CORPORATE SOURCE: Departments of Surgical Oncology, Anderson Cancer  
Center, The University of Texas M. D., Houston, TX,  
77030, USA

SOURCE: Cancer Res. (1999), 59(21), 5412-5416

CODEN: CNREA8; ISSN: 0008-5472

PUBLISHER: AACR Subscription Office

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Increased vascular endothelial growth factor (VEGF) expression is assocd.  
with colon cancer metastases. We hypothesized that inhibition of VEGF  
receptor activity could inhibit colon cancer liver metastases. BALB/c  
mice underwent splenic injection with CT-26 colon cancer cells to generate  
metastases. Mice received daily i.p. injections of vehicle, tyrosine  
kinase inhibitor for Flk-1/KDR (SU5416) or tyrosine kinase inhibitor for  
VEGF, basic fibroblast growth factor, and platelet-derived growth factor  
receptors (SU6668). SU5416 and SU6668 resp. inhibited metastases (48.1%  
and 55.3%), microvessel formation (42.0% and 36.2%), and cell  
proliferation (24.4% and 27.3%) and increased tumor cell (by 2.6- and  
4.3-fold) and endothelial cell (by 18.6- and 81.4-fold) apoptosis (P <  
0.001). VEGF receptor inhibitors increased endothelial cell apoptosis,  
suggesting that VEGF may serve as an endothelial survival factor.

IT 204005-46-9, SU 5416

RL: BAC (Biological activity or effector, except adverse); THU  
(Therapeutic use); BIOL (Biological study); USES (Uses)  
(antiangiogenic therapy targeting VEGF receptor tyrosine  
kinase inhibits liver metastasis of colon cancer and induces tumor and  
endothelial cell apoptosis)

REFERENCE COUNT: 15

REFERENCE(S): (1) Bussolino, F; Trends Biochem Sci 1997, V22, P251  
HCAPLUS  
(2) Dong, Z; J Natl Cancer Inst 1994, V86, P913  
HCAPLUS  
(3) Folkman, J; Nat Med 1995, V1, P27 HCAPLUS

(5) Fong, T; Cancer Res 1999, V59, P99 HCAPLUS  
 (6) Gerber, H; J Biol Chem 1998, V273, P13313 HCAPLUS  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 14 OF 16 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1999:65738 HCAPLUS

DOCUMENT NUMBER: 130:246450

TITLE: SU5416 is a potent and selective inhibitor of the vascular endothelial growth factor receptor (Flk-1/KDR) that inhibits tyrosine kinase catalysis, tumor vascularization, and growth of multiple tumor types

AUTHOR(S): Fong, T. Annie T.; Shawver, Laura K.; Sun, Li; Tang, Cho; App, Harald; Powell, T. Jeff; Kim, Young H.; Schreck, Randall; Wang, Xueyan; Risau, Werner; Ullrich, Axel; Hirth, K. Peter; McMahon, Gerald

CORPORATE SOURCE: SUGEN, Inc., South San Francisco, CA, 94080, USA

SOURCE: Cancer Res. (1999), 59(1), 99-106

CODEN: CNREA8; ISSN: 0008-5472

PUBLISHER: AACR Subscription Office

DOCUMENT TYPE: Journal

LANGUAGE: English

AB SU5416, a novel synthetic compd., is a potent and selective inhibitor of the Flk-1/KDR receptor tyrosine kinase that is presently under evaluation in Phase I clin. studies for the treatment of human cancers. SU5416 was shown to inhibit vascular endothelial growth factor-dependent mitogenesis of human endothelial cells without inhibiting the growth of a variety of tumor cells in vitro. In contrast, systemic administration of SU5416 at nontoxic doses in mice resulted in inhibition of s.c. tumor growth of cells derived from various tissue origins. The antitumor effect of SU5416 was accompanied by the appearance of pale white tumors that were resected from drug-treated animals, supporting the antiangiogenic property of this agent. These findings support that pharmacol. inhibition of the enzymic activity of the vascular endothelial growth factor receptor represents a novel strategy for limiting the growth of a wide variety of tumor types.

IT 204005-46-9, SU 5416

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(SU5416: selective inhibitor of Flk-1/KDR receptor tyrosine kinase, tumor vascularization and growth)

REFERENCE COUNT: 54

REFERENCE(S): (2) Asano, M; Cancer Res 1995, V55, P5296 HCAPLUS  
 (4) Brogi, E; Circulation 1994, V90, P649 HCAPLUS  
 (5) Brown, L; Cancer Res 1993, V53, P4727 HCAPLUS  
 (6) Bussolino, F; Trends Biochem Sci 1997, V22, P251 HCAPLUS  
 (7) Carmeliet, P; Nature (Lond) 1996, V380, P435 HCAPLUS

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 15 OF 16 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1997:640690 HCAPLUS

DOCUMENT NUMBER: 127:314804

TITLE: Assays for KDR/FLK-1 receptor tyrosine kinase inhibitors, and use of the inhibitors for treatment of vasculogenesis- and angiogenesis-related diseases

INVENTOR(S): Hirth, Klaus P.; McMahon, Gerald; Shawver, Laura K.

PATENT ASSIGNEE(S): Sugan, Inc., USA

SOURCE: PCT Int. Appl., 65 pp.

CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9734920	A1	19970925	WO 1997-US3378	19970304
W: AL, AM, AU, AZ, BA, BB, BG, BR, BY, CA, CN, CU, CZ, EE, GE, GH, HU, IL, IS, JP, KG, KP, KR, KZ, LC, LK, LR, LT, LV, MD, MG, MK, MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, TJ, TM, TR, TT, UA, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
AU 9720667	A1	19971010	AU 1997-20667	19970304
PRIORITY APPLN. INFO.:			US 1996-621734	19960321
			WO 1997-US3378	19970304

AB Processes are disclosed for the identification of compds. and pharmaceutical compns. capable of selectively and potently inhibiting KDR/FLK-1 tyrosine kinase signal transduction in order to inhibit vasculogenesis and/or angiogenesis. The invention also relates to compds. and compns. identified using the methods of the invention and the use thereof for the treatment of disease relating to inappropriate vasculogenesis and/or angiogenesis. The invention provides an assay cascade comprised of several "filter steps" of increasing selectivity which identify a limited subset of candidate compds. affecting the VEGF receptor on the mol. level.

IT 3359-49-7, SU 4928 5812-07-7, SU 4312 62540-08-3  
 , SU 5208 91822-51-4, SU 4314 186611-03-0, SU 4932 186611-55-2, SU 4313 204005-46-9, SU 5416  
 RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (KDR/FLK-1 receptor tyrosine kinase inhibitor identification assay, and use of compds. for treatment of vasculogenesis- and angiogenesis-related diseases)

L14 ANSWER 16 OF 16 HCAPLUS COPYRIGHT 2001 ACS  
 ACCESSION NUMBER: 1995:813058 HCAPLUS  
 DOCUMENT NUMBER: 123:208831  
 TITLE: Biologically active 3-substituted oxindole derivatives useful as anti-angiogenic agents  
 INVENTOR(S): Heath, William Francis Heat, Jr.; McDonald, John Hampton III; Brasca, Maria Gabriella; Orzi, Fabrizio; Crugnola, Angelo; Ballinari, Dario; Mariani, Mariangela  
 PATENT ASSIGNEE(S): Pharmacia S.P.A., Italy  
 SOURCE: PCT Int. Appl., 104 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9517181	A1	19950629	WO 1994-EP3664	19941108
W: AU, BY, CA, HU, JP, KR, KZ, NO, PL, RU, UA				



RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE

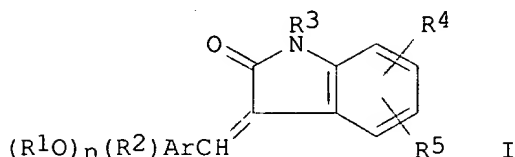
CA 2155098	AA	19950629	CA 1994-2155098	19941108
AU 9480612	A1	19950710	AU 1994-80612	19941108
AU 676958	B2	19970327		
EP 684820	A1	19951206	EP 1994-931583	19941108
EP 684820	B1	20010816		

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, NL, PT, SE

HU 73176	A2	19960628	HU 1995-2761	19941108
JP 08507089	T2	19960730	JP 1994-517121	19941108
AT 204168	E	20010915	AT 1994-931583	19941108
ZA 9410204	A	19951110	ZA 1994-10204	19941212
US 5576330	A	19961119	US 1994-354215	19941212
IL 112010	A1	19981030	IL 1994-112010	19941216
NO 9503146	A	19950810	NO 1995-3146	19950810

PRIORITY APPLN. INFO.: GB 1993-26136 A 19931222  
WO 1994-EP3664 W 19941108

OTHER SOURCE(S): MARPAT 123:208831  
GI



AB Compds. I (Ar = naphthalene, tetralin, quinoline, isoquinoline, indole; n = 0 or an integer of 1 to 3; R<sup>1</sup> = H, C1-6 alkyl, C2-6 alkanoyl; R<sup>2</sup> = H, halogen, C1-6 alkyl, cyano, carboxy, nitro, NHR; R = H, C1-6 alkyl; R<sup>3</sup> = H, C1-6 alkyl; R<sup>4</sup> = H, OH, C1-6 alkoxy, C2-6 alkanoyloxy, carboxy, nitro, NHR; R<sup>5</sup> = H, C1-6 alkyl, halogen) or a pharmaceutically acceptable salt thereof are useful as angiogenesis inhibitors. Products contg. an angiogenesis inhibitor or a pharmaceutically acceptable salt thereof and an antitumor agent are used as a combined prepn. for anticancer therapy. A compn. (for 10,000 tablets) contg. 3-[(3'-hydroxy-2'-tetralyl)methylen]-2-oxindole 250, lactose 800, corn starch 415, talc 30 and Mg stearate 5 g, resp., was formulated.

IT 22813-84-9 55160-03-7 137478-38-7  
137478-39-8 137478-40-1 137479-10-8  
137479-11-9 137479-12-0 137479-13-1  
137479-19-7 137479-20-0 137479-21-1  
137501-13-4 148563-43-3 148563-44-4  
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148563-48-8 148563-49-9 148563-50-2  
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148563-54-6 148563-55-7 148563-56-8  
148563-57-9 148563-58-0 148563-59-1  
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168121-42-4 168121-43-5 168121-44-6  
168121-45-7 168121-46-8 168121-47-9  
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168121-51-5 168121-52-6

RL: BAC (Biological activity or effector, except adverse); THU  
(Therapeutic use); BIOL (Biological study); USES (Uses)  
(oxindole derivs. as anti-angiogenic agents)

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<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

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51	RN	148563-55-7	REGISTRY
52	RN	148563-54-6	REGISTRY
53	RN	148563-53-5	REGISTRY
54	RN	148563-52-4	REGISTRY
55	RN	148563-51-3	REGISTRY
56	RN	148563-50-2	REGISTRY
57	RN	148563-49-9	REGISTRY
58	RN	148563-48-8	REGISTRY
59	RN	148563-47-7	REGISTRY
60	RN	148563-46-6	REGISTRY
61	RN	148563-45-5	REGISTRY
62	RN	148563-44-4	REGISTRY
63	RN	148563-43-3	REGISTRY
64	RN	137501-13-4	REGISTRY
65	RN	137479-21-1	REGISTRY
66	RN	137479-20-0	REGISTRY
67	RN	137479-19-7	REGISTRY
68	RN	137479-13-1	REGISTRY
69	RN	137479-12-0	REGISTRY
70	RN	137479-11-9	REGISTRY
71	RN	137479-10-8	REGISTRY
72	RN	137478-40-1	REGISTRY
73	RN	137478-39-8	REGISTRY

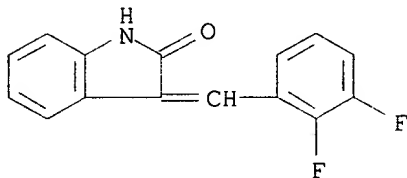
74 RN 137478-38-7 REGISTRY  
 75 RN 91822-51-4 REGISTRY  
 76 RN 62540-08-3 REGISTRY  
 77 RN 55160-03-7 REGISTRY  
 78 RN 22813-84-9 REGISTRY  
 79 RN 5812-07-7 REGISTRY  
 80 RN 3367-88-2 REGISTRY  
 81 RN 3359-49-7 REGISTRY

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, => d ide can l15 1 3 5 10 15 23 26 27 28 30 46 47 64 65 72 75 76 77 78 79 80 81

L15 ANSWER 1 OF 81 REGISTRY COPYRIGHT 2001 ACS  
 RN 342808-38-2 REGISTRY  
 CN 2H-Indol-2-one, 3-[(2,3-difluorophenyl)methylene]-1,3-dihydro- (9CI) (CA  
 INDEX NAME)  
 MF C15 H9 F2 N O  
 SR CA  
 LC STN Files: CA, CAPLUS



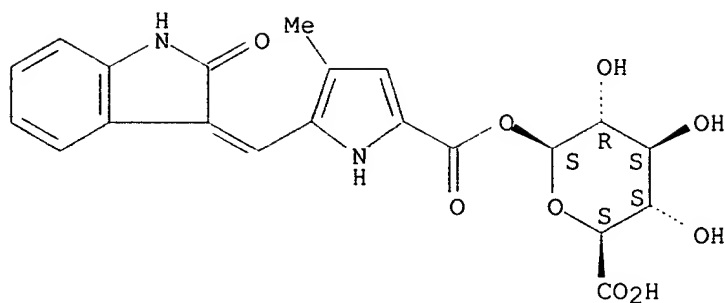
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1967 TO DATE)  
 1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 135:19497

L15 ANSWER 3 OF 81 REGISTRY COPYRIGHT 2001 ACS  
 RN 324047-05-4 REGISTRY  
 CN .beta.-D-Glucopyranuronic acid, 1-[5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-4-methyl-1H-pyrrole-2-carboxylate] (9CI) (CA INDEX NAME)  
 FS STEREOSEARCH  
 MF C21 H20 N2 O9  
 SR CA  
 LC STN Files: CA, CAPLUS, TOXLIT

Absolute stereochemistry.  
 Double bond geometry unknown.



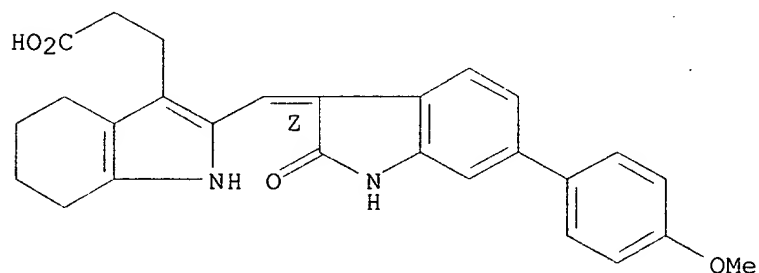
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1967 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 134:141334

L15 ANSWER 5 OF 81 REGISTRY COPYRIGHT 2001 ACS  
RN 288144-36-5 REGISTRY  
CN 1H-Indole-3-propanoic acid, 2-[(Z)-[1,2-dihydro-6-(4-methoxyphenyl)-2-oxo-3H-indol-3-ylidene)methyl]-4,5,6,7-tetrahydro- (9CI) (CA INDEX NAME)  
FS STEREOSEARCH  
MF C27 H26 N2 O4  
SR CA  
LC STN Files: CA, CAPLUS, TOXLIT

Double bond geometry as shown.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

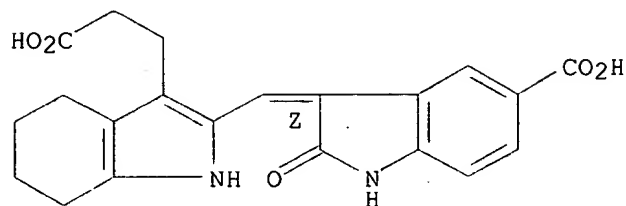
1 REFERENCES IN FILE CA (1967 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 133:159618

L15 ANSWER 10 OF 81 REGISTRY COPYRIGHT 2001 ACS  
RN 288144-31-0 REGISTRY  
CN 1H-Indole-3-propanoic acid, 2-[(Z)-(5-carboxy-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-4,5,6,7-tetrahydro- (9CI) (CA INDEX NAME)  
FS STEREOSEARCH  
MF C21 H20 N2 O5

SR CA  
LC STN Files: CA, CAPLUS, TOXLIT

Double bond geometry as shown.



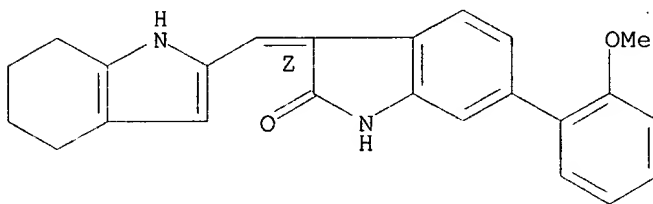
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1967 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 133:159618

L15 ANSWER 15 OF 81 REGISTRY COPYRIGHT 2001 ACS  
RN 288144-26-3 REGISTRY  
CN 2H-Indol-2-one, 1,3-dihydro-6-(2-methoxyphenyl)-3-[(4,5,6,7-tetrahydro-1H-indol-2-yl)methylene]-, (3Z)- (9CI) (CA INDEX NAME)  
FS STEREOSEARCH  
MF C24 H22 N2 O2  
SR CA  
LC STN Files: CA, CAPLUS, TOXLIT

Double bond geometry as shown.



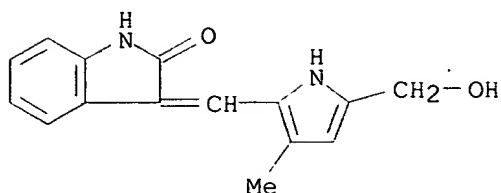
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1967 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 133:159618

L15 ANSWER 23 OF 81 REGISTRY COPYRIGHT 2001 ACS  
RN 280748-41-6 REGISTRY  
CN 2H-Indol-2-one, 1,3-dihydro-3-[[5-(hydroxymethyl)-3-methyl-1H-pyrrol-2-yl)methylene]- (9CI) (CA INDEX NAME)  
OTHER NAMES:  
CN SU 9838  
FS 3D CONCORD  
MF C15 H14 N2 O2

SR CA  
LC STN Files: CA, CAPLUS, TOXLIT



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

3 REFERENCES IN FILE CA (1967 TO DATE)  
1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA  
3 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 135:107247

REFERENCE 2: 134:141334

REFERENCE 3: 133:84238

L15 ANSWER 26 OF 81 REGISTRY COPYRIGHT 2001 ACS

RN 215543-92-3 REGISTRY

CN 1H-Pyrrole-3-propanoic acid, 2-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-4-methyl- (9CI) (CA INDEX NAME)

OTHER NAMES:

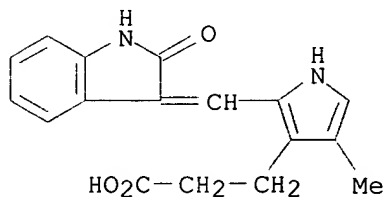
CN SU 5402

FS 3D CONCORD

MF C17 H16 N2 O3

SR CA

LC STN Files: BIOSIS, CA, CAPLUS, DRUGUPDATES, PHAR, TOXLIT, USPATFULL



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

7 REFERENCES IN FILE CA (1967 TO DATE)  
7 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 135:56059

REFERENCE 2: 134:159784

REFERENCE 3: 133:246860

REFERENCE 4: 133:159618  
 REFERENCE 5: 132:166123  
 REFERENCE 6: 132:137242  
 REFERENCE 7: 130:3771

L15 ANSWER 27 OF 81 REGISTRY COPYRIGHT 2001 ACS

RN 204005-46-9 REGISTRY

CN 2H-Indol-2-one, 3-[(3,5-dimethyl-1H-pyrrol-2-yl)methylene]-1,3-dihydro-  
 (9CI) (CA INDEX NAME)

OTHER NAMES:

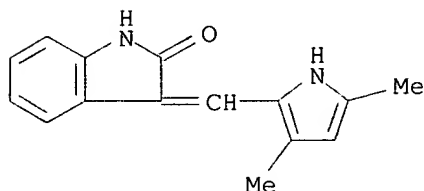
CN SU 5416

FS 3D CONCORD

MF C15 H14 N2 O

SR CA

LC STN Files: BIOSIS, CA, CAPLUS, DRUGPAT, DRUGUPDATES, TOXLIT, USPATFULL



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

37 REFERENCES IN FILE CA (1967 TO DATE)

37 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 135:146994  
 REFERENCE 2: 135:116407  
 REFERENCE 3: 135:107247  
 REFERENCE 4: 135:86674  
 REFERENCE 5: 135:56059  
 REFERENCE 6: 135:2309  
 REFERENCE 7: 134:348057  
 REFERENCE 8: 134:324500  
 REFERENCE 9: 134:316158  
 REFERENCE 10: 134:242681

L15 ANSWER 28 OF 81 REGISTRY COPYRIGHT 2001 ACS

RN 186611-55-2 REGISTRY

CN 2H-Indol-2-one, 1,3-dihydro-3-[[4-(1-methylethyl)phenyl)methylene]- (9CI)  
 (CA INDEX NAME)



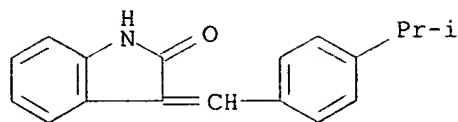
OTHER NAMES:

CN SU 4313

MF C18 H17 N O

SR CA

LC STN Files: CA, CAPLUS, CHEMCATS, TOXLIT, USPATFULL



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

5 REFERENCES IN FILE CA (1967 TO DATE)

5 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 133:234344

REFERENCE 2: 130:237471

REFERENCE 3: 130:237470

REFERENCE 4: 127:314804

REFERENCE 5: 126:139901

L15 ANSWER 30 OF 81 REGISTRY COPYRIGHT 2001 ACS

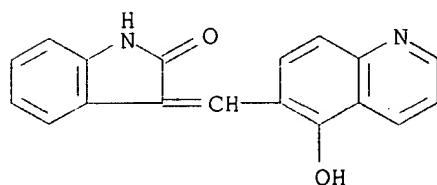
RN 168121-52-6 REGISTRY

CN 2H-Indol-2-one, 1,3-dihydro-3-[(5-hydroxy-6-quinolinyl)methylene]- (9CI)  
(CA INDEX NAME)

MF C18 H12 N2 O2

SR CA

LC STN Files: CA, CAPLUS, TOXLIT, USPATFULL



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1967 TO DATE)

1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

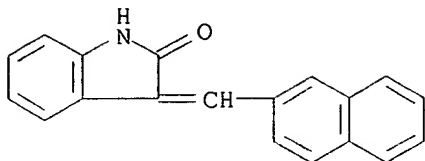
REFERENCE 1: 123:208831

L15 ANSWER 46 OF 81 REGISTRY COPYRIGHT 2001 ACS

RN 149492-57-9 REGISTRY

CN 2H-Indol-2-one, 1,3-dihydro-3-(2-naphthalenylmethylene)- (9CI) (CA INDEX

NAME)  
 MF C19 H13 N O  
 SR CA  
 LC STN Files: CA, CAPLUS, TOXLIT, USPATFULL



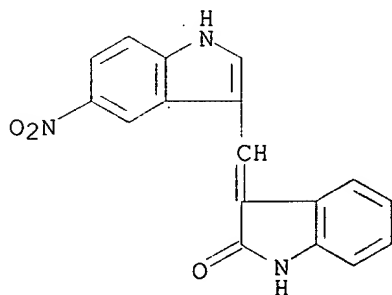
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2 REFERENCES IN FILE CA (1967 TO DATE)  
 2 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 123:208831

REFERENCE 2: 119:108407

L15 ANSWER 47 OF 81 REGISTRY COPYRIGHT 2001 ACS  
 RN 148563-59-1 REGISTRY  
 CN 2H-Indol-2-one, 1,3-dihydro-3-[(5-nitro-1H-indol-3-yl)methylene]- (9CI)  
 (CA INDEX NAME)  
 FS 3D CONCORD  
 MF C17 H11 N3 O3  
 SR CA  
 LC STN Files: CA, CAPLUS, CHEMCATS, TOXLIT, USPATFULL



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

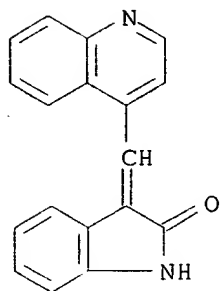
2 REFERENCES IN FILE CA (1967 TO DATE)  
 2 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 123:208831

REFERENCE 2: 119:49225

L15 ANSWER 64 OF 81 REGISTRY COPYRIGHT 2001 ACS  
 RN 137501-13-4 REGISTRY

CN 2H-Indol-2-one, 1,3-dihydro-3-(4-quinolinylmethylene)- (9CI) (CA INDEX  
NAME)  
FS 3D CONCORD  
MF C18 H12 N2 O  
SR CA  
LC STN Files: CA, CAPLUS, TOXLIT, USPATFULL



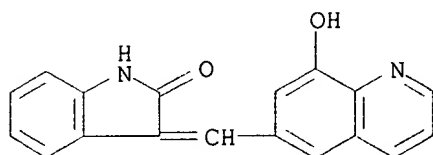
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

2 REFERENCES IN FILE CA (1967 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 123:208831

REFERENCE 2: 115:279621

L15 ANSWER 65 OF 81 REGISTRY COPYRIGHT 2001 ACS  
RN 137479-21-1 REGISTRY  
CN 2H-Indol-2-one, 1,3-dihydro-3-[(8-hydroxy-6-quinolinyl)methylene]- (9CI)  
(CA INDEX NAME)  
MF C18 H12 N2 O2  
SR CA  
LC STN Files: CA, CAPLUS, TOXLIT, USPATFULL



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

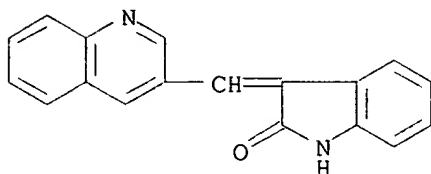
2 REFERENCES IN FILE CA (1967 TO DATE)  
2 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 123:208831

REFERENCE 2: 115:279621

L15 ANSWER 72 OF 81 REGISTRY COPYRIGHT 2001 ACS

RN 137478-40-1 REGISTRY  
 CN 2H-Indol-2-one, 1,3-dihydro-3-(3-quinolinylmethylene)- (9CI) (CA INDEX NAME)  
 MF C18 H12 N2 O  
 SR CA  
 LC STN Files: CA, CAPLUS, TOXLIT, USPATFULL



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

3 REFERENCES IN FILE CA (1967 TO DATE)  
 3 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 123:208831

REFERENCE 2: 119:108407

REFERENCE 3: 115:279621

L15 ANSWER 75 OF 81 REGISTRY COPYRIGHT 2001 ACS

RN 91822-51-4 REGISTRY

CN 2H-Indol-2-one, 1,3-dihydro-3-(1H-pyrrol-2-ylmethylene)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 2-Indolinone, 3-(pyrrol-2-ylmethylene)- (7CI)

OTHER NAMES:

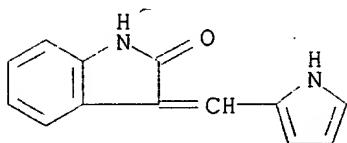
CN HA 12-16

CN SU 4314

FS 3D CONCORD

MF C13 H10 N2 O

LC STN Files: BEILSTEIN\*, CA, CAOLD, CAPLUS, TOXLIT, USPATFULL  
 (\*File contains numerically searchable property data)



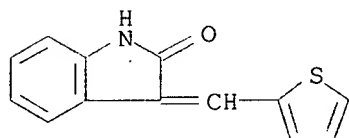
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9 REFERENCES IN FILE CA (1967 TO DATE)  
 9 REFERENCES IN FILE CAPLUS (1967 TO DATE)  
 1 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 133:234344

REFERENCE 2: 132:117566  
 REFERENCE 3: 131:254128  
 REFERENCE 4: 130:237471  
 REFERENCE 5: 130:237470  
 REFERENCE 6: 129:175549  
 REFERENCE 7: 128:204803  
 REFERENCE 8: 127:314804  
 REFERENCE 9: 126:139901

L15 ANSWER 76 OF 81 REGISTRY COPYRIGHT 2001 ACS  
 RN 62540-08-3 REGISTRY  
 CN 2H-Indol-2-one, 1,3-dihydro-3-(2-thienylmethylene)- (9CI) (CA INDEX NAME)  
 OTHER NAMES:  
 CN SU 5208  
 FS 3D CONCORD  
 MF C13 H9 N O S  
 LC STN Files: BEILSTEIN\*, CA, CAPLUS, CHEMCATS, IFICDB, IFIPAT, IFIUDB,  
 TOXLIT, USPATFULL  
 (\*File contains numerically searchable property data)

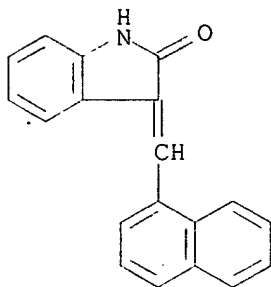


\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

9 REFERENCES IN FILE CA (1967 TO DATE)  
 9 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 130:252240  
 REFERENCE 2: 130:237471  
 REFERENCE 3: 130:237470  
 REFERENCE 4: 129:330650  
 REFERENCE 5: 129:216480  
 REFERENCE 6: 129:175549  
 REFERENCE 7: 127:314804  
 REFERENCE 8: 126:139901  
 REFERENCE 9: 86:171457

L15 ANSWER 77 OF 81 REGISTRY COPYRIGHT 2001 ACS  
 RN 55160-03-7 REGISTRY  
 CN 2H-Indol-2-one, 1,3-dihydro-3-(1-naphthalenylmethylene)- (9CI) (CA INDEX NAME)  
 FS 3D CONCORD  
 MF C19 H13 N O  
 LC STN Files: CA, CAPLUS, CHEMCATS, TOXLIT, USPATFULL



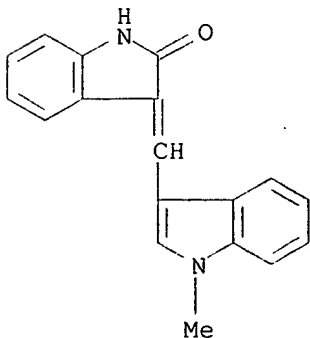
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2 REFERENCES IN FILE CA (1967 TO DATE)  
 2 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 123:208831

REFERENCE 2: 82:139883

L15 ANSWER 78 OF 81 REGISTRY COPYRIGHT 2001 ACS  
 RN 22813-84-9 REGISTRY  
 CN 2H-Indol-2-one, 1,3-dihydro-3-[(1-methyl-1H-indol-3-yl)methylene]- (9CI)  
 (CA INDEX NAME)  
 OTHER CA INDEX NAMES:  
 CN 2-Indolinone, 3-[(1-methylindol-3-yl)methylene]- (8CI)  
 FS 3D CONCORD  
 MF C18 H14 N2 O  
 LC STN Files: BEILSTEIN\*, CA, CAPLUS, TOXLIT, USPATFULL  
 (\*File contains numerically searchable property data)



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

4 REFERENCES IN FILE CA (1967 TO DATE)  
4 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 128:204803

REFERENCE 2: 123:208831

REFERENCE 3: 119:49225

REFERENCE 4: 71:3203

L15 ANSWER 79 OF 81 REGISTRY COPYRIGHT 2001 ACS

RN 5812-07-7 REGISTRY

CN 2H-Indol-2-one, 3-[[4-(dimethylamino)phenyl]methylene]-1,3-dihydro- (9CI)  
(CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 2-Indolinone, 3-[p-(dimethylamino)benzylidene]- (7CI, 8CI)

OTHER NAMES:

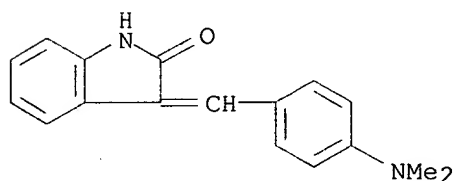
CN 3-(4-Dimethylaminobenzylidenyl)-2-indolinone

CN SU 4312

MF C17 H16 N2 O

LC STN Files: BEILSTEIN\*, BIOSIS, CA, CAOLD, CAPLUS, CHEMCATS, TOXLIT,  
USPATFULL

(\*File contains numerically searchable property data)



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

13 REFERENCES IN FILE CA (1967 TO DATE)  
13 REFERENCES IN FILE CAPLUS (1967 TO DATE)  
1 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 133:234344

REFERENCE 2: 130:348928

REFERENCE 3: 130:252240

REFERENCE 4: 130:237471

REFERENCE 5: 130:237470

REFERENCE 6: 130:149064

REFERENCE 7: 129:330650

REFERENCE 8: 129:175549

REFERENCE 9: 127:314804

REFERENCE 10: 126:139901

L15 ANSWER 80 OF 81 REGISTRY COPYRIGHT 2001 ACS

RN 3367-88-2 REGISTRY

CN 2H-Indol-2-one, 1,3-dihydro-3-(2-pyridinylmethylene)- (9CI) (CA INDEX NAME)

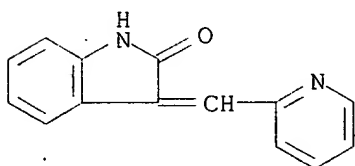
OTHER CA INDEX NAMES:

CN 2-Indolinone, 3-(2-pyridylmethylene)- (7CI, 8CI)

FS 3D CONCORD

MF C14 H10 N2 O

LC STN Files: BEILSTEIN\*, CA, CAOLD, CAPLUS, CHEMCATS, CHEMINFORMRX, IFICDB, IFIPAT, IFIUDB, USPATFULL  
(\*File contains numerically searchable property data)



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

4 REFERENCES IN FILE CA (1967 TO DATE)

4 REFERENCES IN FILE CAPLUS (1967 TO DATE)

3 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 135:19497

REFERENCE 2: 127:190643

REFERENCE 3: 123:169529

REFERENCE 4: 86:171457

L15 ANSWER 81 OF 81 REGISTRY COPYRIGHT 2001 ACS

RN 3359-49-7 REGISTRY

CN 2H-Indol-2-one, 1,3-dihydro-3-(phenylmethylene)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 2-Indolinone, 3-benzylidene- (7CI, 8CI)

CN Oxindole, 3-benzylidene- (6CI)

OTHER NAMES:

CN 1,3-Dihydro-3-(phenylmethylene)-2H-indol-2-one

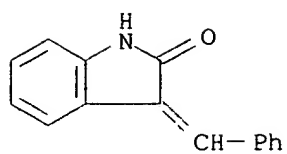
CN 3-Benzylidene-2-indolinone

CN SU 4928

MF C15 H11 N O

LC STN Files: BEILSTEIN\*, CA, CAOLD, CAPLUS, CHEMCATS, IFICDB, IFIPAT, IFIUDB, TOXLIT, USPATFULL  
(\*File contains numerically searchable property data)





\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

18 REFERENCES IN FILE CA (1967 TO DATE)  
18 REFERENCES IN FILE CAPLUS (1967 TO DATE)  
6 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 133:30707  
REFERENCE 2: 130:252240  
REFERENCE 3: 130:237471  
REFERENCE 4: 130:237470  
REFERENCE 5: 129:330650  
REFERENCE 6: 129:175549  
REFERENCE 7: 127:314804  
REFERENCE 8: 126:139901  
REFERENCE 9: 115:8574  
REFERENCE 10: 112:98408

=> fil hcaplus

FILE 'HCAPLUS' ENTERED AT 09:08:11 ON 16 OCT 2001

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FILE COVERS 1947 - 16 Oct 2001 VOL 135 ISS 17

FILE LAST UPDATED: 15 Oct 2001 (20011015/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

This file supports REGISTRY for direct browsing and searching of all substance data from the REGISTRY file. Enter HELP FIRST for more information.

HCAplus now provides online access to patents and literature covered in CA from 1947 to the present. On April 22, 2001, bibliographic information and abstracts were added for over 2.2 million references published in CA from 1947 to 1966.

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L3          STR
L5          4211 SEA FILE=REGISTRY SSS FUL L3
L7          208 SEA FILE=HCAPLUS ABB=ON PLU=ON L5
L11         361 SEA FILE=REGISTRY ABB=ON PLU=ON ANGIOGEN? OR FH1 OR FLK?
L12         17556 SEA FILE=HCAPLUS ABB=ON PLU=ON L11 OR ?ANGIOGEN? OR FH1 OR
          FLK? OR FH(W)1
L13         45 SEA FILE=HCAPLUS ABB=ON PLU=ON L5 AND L12
L14         16 SEA FILE=HCAPLUS ABB=ON PLU=ON L7(L)L12
L16         68 SEA FILE=HCAPLUS ABB=ON PLU=ON L7(L) (INHIBIT? OR ?ANTAGON?)
L17         28 SEA FILE=HCAPLUS ABB=ON PLU=ON L13 AND L16
L18         17 SEA FILE=HCAPLUS ABB=ON PLU=ON L17 NOT L14

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L18 ANSWER 1 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2001:661426 HCAPLUS

DOCUMENT NUMBER: 135:226880

TITLE: Synthesis of Pyrolyllactone-indolinone derivatives as kinase inhibitors

INVENTOR(S): Tang, Peng Cho; Miller, Todd A.; Li, Xiaoyuan; Zhang, Ruofei; Cui, Jingrong; Huang, Ping; Wei, Chung Chen

PATENT ASSIGNEE(S): Sugen, Inc., USA

SOURCE: PCT Int. Appl., 148 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001064681	A2	20010907	WO 2001-US6214	20010228
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
PRIORITY APPLN. INFO.:			US 2000-185536	P 20000228

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\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

AB Title compds. I [R1 = H, alkyl, (hetero)arom. ring, (hetero)aliph. ring, etc.; R2-5 = H, alkyl, (hetero)arom. ring, (hetero)aliph. ring; A, B, D, E = C or N provided that one or two = N and provided that when A, B, D or E = N, no R1 is attached to A, B, D or E; m = 2 - 4; q = 1 - 4] were prepd. Examples include data for over 50 compds. synthesized and over 20 bioassays (data for 4 bioassays provided). For instance, tosylmethyl isocyanide was added to 5,6-dihydro-2H-pyran-2-one (DBU, THF, 0.degree.C, room temp., 2 h.) to give 6,7-dihydro-2H-pyrano[3,4-c]pyrrol-4-one. This intermediate was formylated in the 1-position (DMF, POCl3, DCM, room temp.; 1 h.) followed by condensation of the 1-formyl deriv. with 2-oxo-2,3-dihydro-1H-indole-5-sulfonic acid N-Me amide to yield II. II had IC50 = 0.005 mM for cdk2/cyclin A and IC50 = 6.64 mM for GST-Flk1. Compds. II are used to treat cancer, e.g., squamous cell carcinoma, astrocytoma, Kaposi's sarcoma, etc.

IT 358732-63-5P 358732-64-6P 358732-65-7P  
 358732-66-8P 358732-67-9P 358732-68-0P  
 358732-69-1P 358732-70-4P 358732-71-5P  
 358732-72-6P, 2-Oxo-3-(4-oxo-2,4,6,7-tetrahydropyrano[3,4-c]pyrrol-1-ylmethylene)-2,3-dihydro-1H-indole-5-sulfonic acid amide  
 358732-73-7P 358732-74-8P 358732-75-9P  
 358732-76-0P, N-[2-Oxo-3-(4-oxo-2,4,6,7-tetrahydropyrano[3,4-c]pyrrol-1-ylmethylene)-2,3-dihydro-1H-indol-6-yl]acetamide  
 358732-77-1P 358732-78-2P 358732-79-3P,  
 2-Oxo-3-(4-oxo-2,4,6,7-tetrahydropyrano[3,4-c]pyrrol-1-ylmethylene)-2,3-dihydro-1H-indole-5-carboxylic acid 358732-80-6P,  
 2-Oxo-3-(4-oxo-2,4,6,7-tetrahydropyrano[3,4-c]pyrrol-1-ylmethylene)-2,3-dihydro-1H-indole-6-carboxylic acid 358732-81-7P  
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 358732-88-4P 358732-90-8P 358732-92-0P  
 358732-94-2P 358732-96-4P 358732-98-6P  
 358732-99-7P 358733-00-3P 358733-01-4P,

3-[2-Oxo-3-(4-oxo-2,4,6,7-tetrahydropyrano[3,4-c]pyrrol-1-ylmethylene)-2,3-dihydro-1H-indol-5-yl]benzoic acid 358733-03-6P,  
 3-[2-Oxo-3-(4-oxo-2,4,6,7-tetrahydropyrano[3,4-c]pyrrol-1-ylmethylene)-2,3-dihydro-1H-indol-6-yl]benzoic acid 358733-05-8P

358733-06-9P 358733-07-0P 358733-08-1P

358733-09-2P 358733-10-5P 358733-11-6P

358733-12-7P 358733-13-8P 358733-14-9P

358733-15-0P 358733-16-1P 358733-17-2P

358733-18-3P 358733-19-4P 358733-20-7P

358733-21-8P 358733-23-0P 358733-24-1P

358733-25-2P 358733-26-3P 358733-27-4P

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(drug; synthesis of Pyrolyllactone-indolinone derivs. as kinase inhibitors)

IT 150977-45-0, Flk-1 receptor tyrosine kinase

RL: BPR (Biological process); BIOL (Biological study); PROC (Process)

(synthesis of Pyrolyllactone-indolinone derivs. as kinase inhibitors)

L18 ANSWER 2 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2001:617993 HCAPLUS

DOCUMENT NUMBER: 135:195497

TITLE: Preparation of pyrrole substituted 2-indolinone protein kinase inhibitors for treatment of cancer

INVENTOR(S): Tang, Peng Cho; Miller, Todd; Li, Xiaoyuan; Sun, Li; Wei, Chung Chen; Shirazian; Shahrzad; Liang, Congxin; Vojkovsky, Tomas; Nematalla, Asaad S.

PATENT ASSIGNEE(S): Sugan, Inc., USA

SOURCE: PCT Int. Appl., 225 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

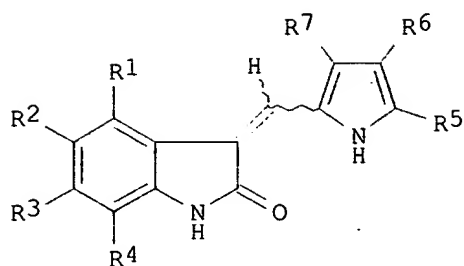
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001060814	A2	20010823	WO 2001-US4813	20010215
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			

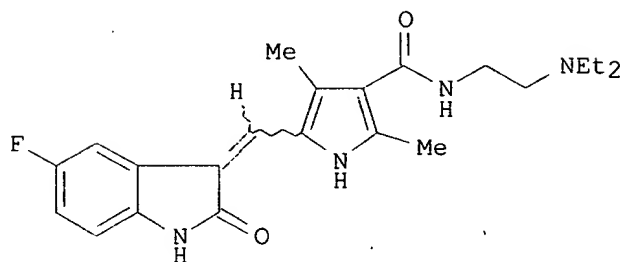
PRIORITY APPLN. INFO.: US 2000-182710 P 20000215  
 US 2000-216422 P 20000706  
 US 2000-243532 P 20001027

OTHER SOURCE(S): MARPAT 135:195497

GI



I



II

AB The title compds. (I) [wherein R1 = H, halo, (cyclo)alkyl, (hetero)aryl, heteroalicyclic, OH, alkoxy, acyl, (un)substituted amino or carbamoyl, etc.; R2 = H, halo, alkyl, trihalomethyl, OH, alkoxy, CN, (hetero)aryl, (un)substituted amino, acyl(amino), or sulfamoyl, etc.; R3 = H, halo, alkyl, trihalomethyl, OH, alkoxy, (hetero)aryl, (un)substituted acyl, (acyl)amino, sulfamoyl, or alkylsulfonyl, etc.; R4 = H, halo, alkyl, OH, alkoxy, or (un)substituted amino; R5 and R6 = independently H, alkyl, or acyl; R7 = H, alkyl, (hetero)aryl, or acyl; and their pharmaceutically acceptable salts] were prepd. as protein kinase modulators for the treatment of cellular disorders such as cancer. For example, 5-fluoro-1,3-dihydroindol-2-one was condensed with 5-formyl-2,4-dimethyl-1H-pyrrole-3-carboxylic acid (2-diethylaminoethyl)amide to give II (55%). II exhibited comparable activity against Flk-1 and PDGFR.beta. and inhibited PDGF-dependent receptor phosphorylation in cells with an IC50 value of approx. 0.03 .mu.M. In efficacy expts. against various cancers in mice, II was well tolerated at 80 mg/kg/day, even when dosed continuously for more than 100 days.

IT 356068-93-4P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation) (intermediate; prepn. of pyrrole substituted 2-indolinone protein kinase inhibitors by condensation of dihydroindolones with formylpyrroles for treatment of cancer and other diseases)

IT 326914-13-0P

RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (prepn. of pyrrole substituted 2-indolinone protein kinase inhibitors by condensation of dihydroindolones with formylpyrroles for treatment of cancer and other diseases)

IT 280748-39-2P 280748-40-5P 326914-09-4P  
326914-10-7P 326914-17-4P 326914-19-6P  
342641-16-1P 342641-17-2P 342641-18-3P  
342641-19-4P 342641-20-7P 342641-21-8P  
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342641-25-2P 342641-26-3P 342641-27-4P  
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 356069-73-3P 356069-74-4P 356069-75-5P  
 356069-76-6P 356069-77-7P

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of pyrrole substituted 2-indolinone protein kinase

**inhibitors** by condensation of dihydroindolones with  
formylpyrroles for treatment of cancer and other diseases)

IT 342641-53-6 356069-06-2 356069-08-4  
356069-10-8 356069-14-2

RL: RCT (Reactant)

(reactant; prepn. of pyrrole substituted 2-indolinone protein kinase  
**inhibitors** by condensation of dihydroindolones with  
formylpyrroles for treatment of cancer and other diseases)

L18 ANSWER 3 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2001:380529 HCAPLUS

DOCUMENT NUMBER: 134:366685

TITLE: Preparation and use of 4-(benzyloxy)phenylalkanoic  
acid plasminogen activator inhibitor antagonists

INVENTOR(S): Madison, Edwin L.; Brunck, Terence K.; Semple, Joseph  
Edward; Lim-Wilby, Marguerita; Pryor, Kent E.; Lewis,  
Ronald D., II; Duncan, David F.; Lawrence, C. Maxwell

PATENT ASSIGNEE(S): Corvas International, Inc., USA

SOURCE: PCT Int. Appl., 261 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

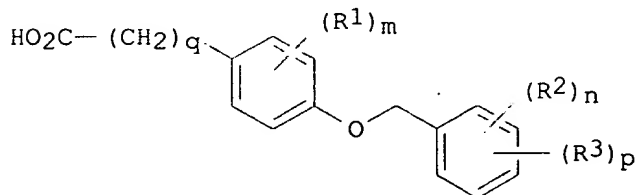
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001036351	A2	20010525	WO 2000-US31824	20001117
W:	AE, AG, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
PRIORITY APPLN. INFO.:			US 1999-444172	A2 19991119
			US 2000-185564	P 20000228
			US 2000-580535	A2 20000526

OTHER SOURCE(S): MARPAT 134:366685  
GI



AB Title compds. (I) [wherein R1 = independently (pseudo)halo, (halo)alkoxy, aryl, or (halo)alkyl; m = 0-4; q = 0-6; when n = 1-5 and p = 0, R2 and R3 = independently NO2, CN, (hetero)arylcarbonyl, haloalkylthio, (pseudo)halo, (cyclo)alkyl, (hetero)aryl, (cyclo)alkoxy, (hetero)aryloxy, haloalkyl, or haloalkoxy; or when n and p = 1, R2 and R3 together with the

C atoms to which they are attached form a (hetero)arom. ring or (hetero)cyclic ring] and pharmaceutical compns. were prepd. as plasminogen activator inhibitor (PAI) antagonists for treatment of thrombotic disorders, unstable angina, cancer, and hemostatic disorders. In particular, methods of antagonizing PAI with substituted and unsubstituted aryl and heteroaryl ethers and thioethers, benzils, benzyl ethers, benzoate esters, sulfones, and benzophenones are provided. For example, a suspension of 3,5-diiodo-4-hydroxybenzoic acid, 3-bromobenzyl bromide, and Cs<sub>2</sub>CO<sub>3</sub> in DMF was stirred for 15 h at room temp. to give 4-(3-bromobenzyloxy)-3,5-diiodobenzoic acid 3-bromobenzyl ester (84%). The IC<sub>50</sub> for PAI antagonist activity for each of the disclosed compds. was measured, and most exhibited IC<sub>50</sub> < 100 .mu.M.

IT 340314-10-5P, 3-(4-Hydroxy-3,5-diiodobenzylidene)-1,3-dihydroindol-2-one

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. and use of 4-(benzyloxy)phenylalkanoic acid plasminogen activator **inhibitor antagonists** for treatment of thrombotic disorders, unstable angina, cancer, and hemostatic disorders)

L18 ANSWER 4 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2001:240686 HCAPLUS

DOCUMENT NUMBER: 135:2309

TITLE: Inhibition of vascular endothelial growth factor receptor signaling leads to reversal of tumor resistance to radiotherapy

AUTHOR(S): Geng, Ling; Donnelly, Edwin; McMahon, Gerald; Lin, P. Charles; Sierra-Rivera, Elaine; Oshinka, Halina; Hallahan, Dennis E.

CORPORATE SOURCE: Departments of Radiation Oncology, Vanderbilt University School of Medicine, Vanderbilt University, Nashville, TN, 37232, USA

SOURCE: Cancer Res. (2001), 61(6), 2413-2419

CODEN: CNREA8; ISSN: 0008-5472

PUBLISHER: American Association for Cancer Research

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Certain refractory neoplasms, such as glioblastoma multiforme (GBM) and melanoma, demonstrate a resistant tumor phenotype in vivo. We obsd. that these refractory tumor models (GBM and melanoma) contain blood vessels that are relatively resistant to radiotherapy. To det. whether the vascular endothelial growth factor receptor-2 (Flk-1/KDR) may be a therapeutic target to improve the effects of radiotherapy, we used the sol. extracellular component of Flk-1 (ExFlk), which blocks vascular endothelial growth factor binding to Flk-1 receptor expressed on the tumor endothelium. Both sFlk-1 and the Flk-1-specific inhibitor SU5416 eliminated the resistance phenotype in GBM and melanoma microvasculature as detd. by both the vascular window and Doppler blood flow methods. Human microendothelial cells and human umbilical vein endothelial cells showed minimal radiation-induced apoptosis. The Flk-1 antagonists sFlk-1 and SU5416 reverted these cell models to apoptosis-prone phenotype. Flk-1 antagonists also reverted GBM and melanoma tumor models to radiation-sensitive phenotype after treatment with 3 Gy. These findings demonstrate that the tumor microenvironment including the survival of tumor-assocd. endothelial cells contributes to tumor blood vessel resistance to therapy.



IT 204005-46-9, SU 5416

RL: BAC (Biological activity or effector, except adverse); THU  
(Therapeutic use); BIOL (Biological study); USES (Uses)  
(tumor vascularization inhibitor; inhibition of  
VEGF receptor signaling enhances radiation-induced response in tumor  
blood vessels)

REFERENCE COUNT: 39

REFERENCE(S): (2) Advani, S; Gene Ther 1998, V5, P160 HCAPLUS  
(9) Fong, T; Cancer Res 1999, V59, P99 HCAPLUS  
(10) Gerber, H; J Biol Chem 1998, V273, P13313 HCAPLUS  
(11) Goldman, C; Proc Natl Acad Sci USA 1998, V95,  
P8795 HCAPLUS  
(12) Gorski, D; Cancer Res 1999, V59, P3374 HCAPLUS  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L18 ANSWER 5 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:688215 HCAPLUS

DOCUMENT NUMBER: 133:252306

TITLE: Preparation of indolinones as protein kinase  
inhibitors.

INVENTOR(S): Tang, Peng Cho; Sun, Li; McMahon, Gerald; Miller, Todd  
Anthony; Shirazian, Shahrzad; Wei, Chung Chen; Harris,  
G. Davis; Xiaoyuan, Li; Liang, Congxin

PATENT ASSIGNEE(S): Sugan, Inc., USA

SOURCE: PCT Int. Appl., 245 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

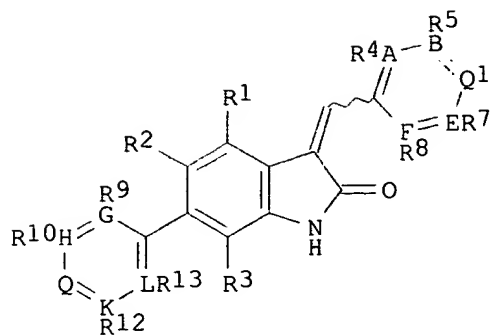
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000056709	A1	20000928	WO 2000-US7704	20000322
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
PRIORITY APPLN. INFO.:			US 1999-125945	P 19990324
			US 1999-127863	P 19990405
			US 1999-131192	P 19990426
			US 1999-132243	P 19990503

OTHER SOURCE(S): MARPAT 133:252306

GI



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AB Title compds., e.g. [I; m, n = 0, 1; Q = (JR11)m; Q1 = (DR6)n; when n = 1, then A, B, D, E, F = C, N; .ltoreq.3 of A, B, D, E, F = N; when m = 1, then G, H, J, K, L = C, N; .gtoreq.1 and .ltoreq.3 of G, H, J, K, L = N; when n = 0, then A = C, N, B, F = C, N, NH, O, S; E = C, N, O, S; when m = O, then G = C, N, H, K, l = C, N, NH, O, S; R1-R13 = H, alkyl, trihaloalkyl, cycloalkyl, alkenyl, alkynyl, aryl, heteroaryl, OH, alkoxy, SH, alkylthiol, aryloxy, amino, etc.; R4R5 or R5R6 or R6R7 or R7R8 = atoms to form a 5-6 membered (hetero)aryl ring; with addnl. provisos], were prepd. Thus, 6-pyridin-3-yl-1,3-dihydroindol-2-one (prepn. given), 4-methoxy-3-thien-2-ylbenzaldehyde, and piperidine were refluxed overnight in EtOH to give 15% 3-(4-methoxy-3-thien-2-ylbenzylidene)-6-pyridin-3-yl-1,3-dihydroindol-2-one. Tested title compds. inhibited HER2 kinase with IC50 = 16.4 .mu.M to .gtoreq.100 .mu.M.

IT 295798-91-3P 295798-95-7P

RL: BAC (Biological activity or effector, except adverse); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
(prepn. of indolinones as protein kinase inhibitors)

IT 251356-74-8P 295798-93-5P 295798-97-9P  
295798-99-1P 295799-27-8P 295799-29-0P  
295799-31-4P 295799-33-6P 295799-35-8P  
295799-37-0P 295799-39-2P 295799-41-6P  
295799-43-8P 295799-47-2P 295799-51-8P  
295799-97-2P

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of indolinones as protein kinase inhibitors)

IT 258828-94-3 258829-04-8 258829-17-3  
258829-24-2 295800-19-0 295800-20-3  
295800-21-4 295800-22-5 295800-23-6  
295800-24-7

RL: RCT (Reactant)

(prepn. of indolinones as protein kinase inhibitors)

IT 251356-61-3P 251356-63-5P 251356-65-7P  
251356-66-8P 251356-67-9P 251356-68-0P  
258829-01-5P 295800-04-3P 295800-06-5P  
295800-07-6P 295800-08-7P 295800-09-8P  
295800-25-8P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation)

(prepn. of indolinones as protein kinase inhibitors)

REFERENCE COUNT: 31

REFERENCE(S): (1) Andreani; DATABASE CHEMABS HCAPLUS

- (2) Andreani; EUR J MED CHEM 1990, V25(2), P187 HCAPLUS  
 (3) Decodts; DATABASE CHEMABS HCAPLUS.  
 (4) Decodts; EUR J MED CHEM - CHIM THER 1983, V18(2), P107 HCAPLUS  
 (5) Erba Carlo Spa; WO 9113055 A 1991 HCAPLUS  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L18 ANSWER 6 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:622463 HCAPLUS  
 DOCUMENT NUMBER: 133:217719  
 TITLE: 3-(Cyclohexanoheteroarylidenyl)-2-indolinone protein tyrosine kinase inhibitors, and their therapeutic use  
 INVENTOR(S): Tang, Peng Cho; Sun, Li; McMahon, Gerald; Blake, Robert A.  
 PATENT ASSIGNEE(S): Sugan, Inc., USA  
 SOURCE: U.S., 61 pp., Cont. -in-part of U.S. Ser. No. 99,842.  
 CODEN: USXXAM  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 2  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6114371	A	20000905	US 1998-190970	19981112
US 6130238	A	20001010	US 1998-99842	19980619
PRIORITY APPLN. INFO.:			US 1997-50977	P 19970620
			US 1997-59384	P 19970919
			US 1998-99842	A2 19980619
			US 1997-59544	P 19970919

OTHER SOURCE(S): CASREACT 133:217719; MARPAT 133:217719

AB 3-(Cyclohexano-heteroarylidenyl)-2-indolinone compds., and physiol. acceptable salts and prodrugs thereof, are disclosed which are expected to modulate the activity of protein tyrosine kinases and therefore to be useful in the prevention and treatment of protein tyrosine kinase-related cellular disorders (cancer, arthritis, restenosis, etc.).

IT 215543-90-1 215543-91-2 245035-88-5  
 245035-93-2 245035-96-5 245036-00-4  
 245036-07-1 245036-08-2 245036-21-9  
 245036-22-0 290821-14-6 290821-15-7  
 290821-16-8 290821-17-9 290821-18-0  
 290821-19-1 290821-20-4

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (cyclohexanoheteroarylidenyl indolinone protein tyrosine kinase inhibitors, and therapeutic use)

IT 150977-45-0, Flk-1 receptor tyrosine kinase

RL: BPR (Biological process); BIOL (Biological study); PROC (Process)  
 (cyclohexanoheteroarylidenyl indolinone protein tyrosine kinase inhibitors, and therapeutic use)

REFERENCE COUNT: 38

REFERENCE(S): (1) Akbasak; J Neurol Sci 1992, V111, P119 HCAPLUS  
 (2) Andreani; Eur J Med Chem 1997, V32, P919 HCAPLUS  
 (3) Anon; WO 9115495 1991 HCAPLUS  
 (4) Anon; WO 9220642 1992 HCAPLUS  
 (5) Anon; WO 9221660 1992 HCAPLUS  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L18 ANSWER 7 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:509389 HCAPLUS

DOCUMENT NUMBER: 134:216785

TITLE: New anilinophthalazines as potent and orally well absorbed inhibitors of the VEGF receptor tyrosine kinases useful as antagonists of tumor-driven **angiogenesis**. [Erratum to document cited in CA133:99079]

AUTHOR(S): Bold, Guido; Altmann, Karl-Heinz; Frei, Joerg; Lang, Marc; Manley, Paul W.; Traxler, Peter; Wietfeld, Bernhard; Brueggen, Josef; Buchdunger, Elisabeth; Cozens, Robert; Ferrari, Stefano; Furet, Pascal; Hofmann, Francesco; Martiny-Baron, Georg; Mestan, Juergen; Roesel, Johannes; Sills, Matthew; Stover, David; Acemoglu, Figan; Boss, Eugen; Emmenegger, Rene; Laesser, Laurent; Masso, Elvira; Roth, Rosemarie; Schlachter, Christian; Vetterli, Werner; Wyss, Dominique; Wood, Jeanette M.

CORPORATE SOURCE: Oncology Research and Process Research, NOVARTIS Pharma AG, Basel, CH-4002, Switz.

SOURCE: J. Med. Chem. (2000), 43(16), 3200

CODEN: JMCMAR; ISSN: 0022-2623

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

AB On page 2316, in Table 3, the unit for cmax; the concn. should be given as [.mu.M]. The correct version of Table 3 is given.

IT 204005-46-9, SU 5416

RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(prepn. of anilinophthalazines as **inhibitors** of VEGF receptor tyrosine kinase (Erratum))

L18 ANSWER 8 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:500184 HCAPLUS

DOCUMENT NUMBER: 133:234344

TITLE: DoMCoSAR: A Novel Approach for Establishing the Docking Mode That Is Consistent with the Structure-Activity Relationship. Application to HIV-1 Protease Inhibitors and VEGF Receptor Tyrosine Kinase Inhibitors

AUTHOR(S): Vieth, Michal; Cummins, David J.

CORPORATE SOURCE: Lilly Research Laboratories, Eli Lilly and Company, Indianapolis, IN, 46285, USA

SOURCE: J. Med. Chem. (2000), 43(16), 3020-3032

CODEN: JMCMAR; ISSN: 0022-2623

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

AB DoMCoSAR is a novel approach for statistically detg. the docking mode that is consistent with a structure-activity relationship. The approach establishes the binding mode for the compds. in a chem. series with the assumption that all mols. exhibit the same binding mode. It involves three stages. In the first stage all mols. that belong to a given chem. series are docked to the active site of the protein target. The only bias used in the docking at this stage involves the location of the protein binding site. Coordinates of the common substructure (CS) that results from the unbiased docking are then clustered to establish the major

substructure docking modes. In the second stage all mols. are docked to the major docking modes (MDMs) with constraints based on the common substructure. The third stage generates, for the major docking modes, interaction-based descriptors that include electrostatic, VDW, strain, and solvation contributions. The problem of docking mode evaluation is now reduced to the question of which descriptor set is more predictive. To establish a quant. comparison of the descriptor sets assocd. with the major docking modes, we use 50 instances of random 4-fold cross-validation. For each 4-fold cross-validation the predictive squared correlation coeff. (R<sup>2</sup>) is computed. T-Tests are applied to establish significance of the differences in mean R for one docking mode vs. another. We test the methodol. on two test cases: HIV-1 protease inhibitors (Holloway et al. J. Med. Chem. 1995, 38, 305-317) and vascular endothelial growth factor (VEGF) receptor tyrosine kinase oxoindoles (Sun et al. J. Med. Chem. 1998, 41, 2588-2603). For both test cases there is statistically significant preference for the binding mode consistent with the x-ray structure. The appeal of this methodol. is that researchers gain the objectivity of statistical justification for the selected docking mode. The methodol. is relatively insensitive to subtle variations of the protein structure that include, but are not limited to, side chain and small backbone rearrangement during binding. In addn., predictive models that result from the approach can be used to further optimize chem. series.

IT 5812-07-7 15966-93-5 40526-64-5  
 55160-02-6 64259-01-4 64259-05-8  
 76086-99-2 91822-51-4 108402-27-3  
 186610-88-8 186610-89-9 186610-90-2  
 186610-94-6 186610-96-8 186611-10-9  
 186611-25-6 186611-27-8 186611-28-9  
 186611-29-0 186611-37-0 186611-38-1  
 186611-40-5 186611-45-0 186611-47-2  
 186611-48-3 186611-53-0 186611-55-2  
 186611-65-4 204005-46-9 204005-54-9  
 293302-08-6 293302-09-7 293302-10-0  
 293302-11-1 293302-12-2 293302-13-3  
 293302-14-4 293302-15-5 293302-16-6  
 293302-17-7 293302-18-8 293302-19-9  
 293302-20-2 293302-22-4 293302-23-5  
 293302-24-6 293302-25-7 293302-26-8  
 293302-27-9 293302-28-0

RL: BPR (Biological process); PRP (Properties); THU (Therapeutic use);  
 BIOL (Biological study); PROC (Process); USES (Uses)  
 (VEGF kinase-inhibitor; DoMCoSAR - novel approach for  
 establishing docking mode that is consistent with structure-activity  
 relationship with application to HIV-1 protease inhibitors  
 and VEGF receptor tyrosine kinase inhibitors)

IT 150977-45-0, Flk-1/KDR VEGF receptor tyrosine kinase  
 RL: BAC (Biological activity or effector, except adverse); BPR (Biological  
 process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);  
 USES (Uses)  
 (inhibitors; DoMCoSAR - novel approach for establishing docking mode  
 that is consistent with structure-activity relationship with  
 application to HIV-1 protease inhibitors and VEGF receptor tyrosine  
 kinase inhibitors)

REFERENCE COUNT: 45

REFERENCE(S): (1) Altschul, S; Nucleic Acid Res 1997, V25, P3389  
 HCAPLUS  
 (3) Brooks, B; J Comput Chem 1983, V4, P187 HCAPLUS  
 (5) Brown, R; J Chem Inf Comput Sci 1997, V37, P1

## HCAPLUS

(7) Charifson, P; J Med Chem 1999, V42, P5100 HCAPLUS  
 (10) Gallop, M; J Med Chem 1994, V37, P1233 HCAPLUS  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L18 ANSWER 9 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:359936 HCAPLUS

DOCUMENT NUMBER: 133:99079

TITLE: New Anilinophthalazines as Potent and Orally Well  
 Absorbed Inhibitors of the VEGF Receptor Tyrosine  
 Kinases Useful as Antagonists of Tumor-Driven  
**Angiogenesis**

AUTHOR(S): Bold, Guido; Altmann, Karl-Heinz; Frei, Joerg; Lang,  
 Marc; Manley, Paul W.; Traxler, Peter; Wietfeld,  
 Bernhard; Brueggel, Josef; Buchdunger, Elisabeth;  
 Cozens, Robert; Ferrari, Stefano; Furet, Pascal;  
 Hofmann, Francesco; Martiny-Baron, Georg; Mestan,  
 Juergen; Roesel, Johannes; Sills, Matthew; Stover,  
 David; Acemoglu, Figan; Boss, Eugen; Emmenegger, Rene;  
 Laesser, Laurent; Masso, Elvira; Roth, Rosemarie;  
 Schlachter, Christian; Vetterli, Werner; Wyss,  
 Dominique; Wood, Jeanette M.

CORPORATE SOURCE: Oncology Research and Process Research, NOVARTIS  
 Pharma AG, Basel, CH-4002, Switz.

SOURCE: J. Med. Chem. (2000), 43(12), 2310-2323  
 CODEN: JMCMAR; ISSN: 0022-2623

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The sprouting of new blood vessels, or **angiogenesis**, is  
 necessary for any solid tumor to grow large enough to cause  
 life-threatening disease. Vascular endothelial growth factor (VEGF) is  
 one of the key promoters of tumor induced **angiogenesis**. VEGF  
 receptors, the tyrosine kinases Flt-1 and KDR, are expressed on vascular  
 endothelial cells and initiate **angiogenesis** upon activation by  
 VEGF. 1-Anilino-(4-pyridylmethyl)-phthalazines, such as CGP 79787D (or  
 PTK787 / ZK222584), reversibly inhibit Flt-1 and KDR with IC50 values <  
 0.1 .mu.M. CGP 79787D also blocks the VEGF-induced receptor  
 autophosphorylation in CHO cells ectopically expressing the KDR receptor  
 (ED50 = 34 nM). Modification of the 1-anilino moiety afforded derivs.  
 with higher selectivity for the VEGF receptor tyrosine kinases Flt-1 and  
 KDR compared to the related receptor tyrosine kinases PDGF-R and c-Kit.  
 Since these 1-anilino-(4-pyridylmethyl)phthalazines are orally well  
 absorbed, these compds. qualify for further profiling and as candidates  
 for clin. evaluation.

IT 204005-46-9, SU 5416

RL: BAC (Biological activity or effector, except adverse); BPR (Biological  
 process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);  
 USES (Uses)

(prepn. of anilinophthalazines as **inhibitors** of VEGF receptor  
 tyrosine kinase)

REFERENCE COUNT: 51

REFERENCE(S): (1) Andersen, L; Acta Chem Scand, Ser B 1988, VB42,  
 P492 HCAPLUS  
 (2) Augustin, H; Trends Pharmacol Sci 1998, V19, P216  
 HCAPLUS  
 (3) Bergers, G; Science 1999, V284, P808 HCAPLUS  
 (4) Bold, G; WO 9835958 A1 1998 HCAPLUS  
 (5) Breier, G; Trends Cell Biol 1996, V6, P454 HCAPLUS

## ALL CITATIONS AVAILABLE IN THE RE FORMAT

L18 ANSWER 10 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1999:757553 HCAPLUS

DOCUMENT NUMBER: 132:137242

TITLE: Design, Synthesis, and Evaluations of Substituted  
3-[(3- or 4-Carboxyethylpyrrol-2-yl)methylidenyl]indolin-2-ones as Inhibitors of VEGF, FGF, and PDGF Receptor Tyrosine Kinases

AUTHOR(S): Sun, Li; Tran, Ngoc; Liang, Congxin; Tang, Flora; Rice, Audie; Schreck, Randall; Waltz, Kara; Shawver, Laura K.; McMahon, Gerald; Tang, Cho

CORPORATE SOURCE: SUGEN Inc., South San Francisco, CA, 94080-4811, USA

SOURCE: J. Med. Chem. (1999), 42(25), 5120-5130

CODEN: JMCMAR; ISSN: 0022-2623

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Receptor tyrosine kinases (RTKs) were implicated as therapeutic targets for the treatment of human diseases including cancers, inflammatory diseases, cardiovascular diseases including arterial restenosis, and fibrotic diseases of the lung, liver, and kidney. Three classes of 3-substituted 2-indolinones contg. propanoic acid functionality attached to the pyrrole ring at the C-3 position of the core were identified as catalytic inhibitors of the vascular endothelial growth factor (VEGF), fibroblast growth factor (FGF), and platelet-derived growth factor (PDGF) RTKs. Some of the compds. were found to inhibit the tyrosine kinase activity assocd. with isolated vascular endothelial growth factor receptor 2 (VEGF-R2) [fetal liver tyrosine kinase 1 (Flk-1)/kinase insert domain-contg. receptor (KDR)], fibroblast growth factor receptor (FGF-R), and platelet-derived growth factor receptor (PDGF-R) tyrosine kinase with IC50 values at nanomolar level. Thus, SU 5402 [5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-4-methyl-1H-Pyrrole-3-propanoic acid] showed inhibition against VEGF-R2 (Flk-1/KDR) and FGF-R1 tyrosine kinase activity with IC50 values of 20 and 30 nM, resp., while 5-[(1,2-dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrole-3-propanoic acid inhibited the PDGF-R tyrosine kinase activity with IC50 value of 10 nM. Structural models and structure-activity relationship anal. of these compds. for the target receptors are discussed. The cellular activities of these compds. were profiled using cellular proliferation assays as measured by bromodeoxyuridine (BrdU) incorporation. Specific and potent inhibition of cell growth was obsd. for some of these compds. These data provide evidence that these compds. can be used to inhibit the function of these target receptors.

IT 256657-46-2 256657-47-3 256657-51-9

RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study)

(prepn. and evaluation of [(dihydrooxoindolylidene)methyl]pyrrolepropanoic acid as tyrosine kinase inhibitors)

IT 186611-14-3P, 5-[(1,2-Dihydro-2-oxo-3H-indol-3-ylidene)methyl]-4-methyl-1H-Pyrrole-3-propanoic acid 245036-27-5P, 5-[(1,2-Dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-Pyrrole-3-propanoic acid 251356-17-9P, 5-[(1,2-Dihydro-6-methoxy-2-oxo-3H-indol-3-ylidene)methyl]-4-methyl-1H-Pyrrole-3-propanoic acid 251356-26-0P 251356-27-1P 251356-32-8P, 5-[(5-Bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-Pyrrole-3-propanoic acid 251356-38-4P, 5-[(1,2-Dihydro-6-methoxy-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-Pyrrole-3-propanoic acid 251356-40-8P, 5-[(1,2-Dihydro-6-(3-methoxyphenyl)-2-oxo-3H-indol-3-

ylidene)methyl]-2,4-dimethyl-1H-Pyrrole-3-propanoic acid  
**251356-42-0P**, 5-[[[1,2-Dihydro-6-(3-methoxyphenyl)-2-oxo-3H-indol-3-ylidene)methyl]-4-methyl-1H-Pyrrole-3-propanoic acid **251356-45-3P**  
 , 5-[[[1,2-Dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-pyrrole-3-propanoic acid **251356-46-4P**, 5-[[[1,2-Dihydro-2-oxo-6-phenyl-3H-indol-3-ylidene)methyl]-4-methyl-1H-Pyrrole-3-propanoic acid **251356-47-5P**, 5-[[[1,2-Dihydro-6-(4-methoxyphenyl)-2-oxo-3H-indol-3-ylidene)methyl]-4-methyl-1H-Pyrrole-3-propanoic acid **251356-48-6P**  
 , 5-[[[1,2-Dihydro-6-(4-methoxyphenyl)-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-Pyrrole-3-propanoic acid **251356-49-7P**,  
 5-[[[1,2-Dihydro-6-(2-methoxyphenyl)-2-oxo-3H-indol-3-ylidene)methyl]-4-methyl-1H-Pyrrole-3-propanoic acid **251356-50-0P**,  
 5-[[[1,2-Dihydro-6-(2-methoxyphenyl)-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-1H-Pyrrole-3-propanoic acid **256657-49-5P**  
**256657-64-4P 256657-65-5P**

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)  
 (prepn. and evaluation of [(dihydrooxoindolylidene)methyl]pyrrolepropanoic acid as tyrosine kinase **inhibitors**)

IT **204005-46-9**, SU 5416 **215543-92-3**

RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (prepn. and evaluation of [(dihydrooxoindolylidene)methyl]pyrrolepropanoic acid as tyrosine kinase **inhibitors**)

REFERENCE COUNT: 15

REFERENCE(S): (1) Cantley, L; Cell 1991, V64, P281 HCAPLUS  
 (3) Hanks, S; Methods Enzymol 1991, V200, P38 HCAPLUS  
 (4) Kolibaba, K; Biochim Biophys Acta 1997, V1333, PF217 HCAPLUS  
 (5) Merenmies, J; Cell Growth Differ 1997, V8, P3 HCAPLUS  
 (6) Mohammadi, M; EMBO J 1998, V17(20), P5896 HCAPLUS  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L18 ANSWER 11 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1999:166598 HCAPLUS

DOCUMENT NUMBER: 130:209599

TITLE: Preparation of benzylidene-1,3-dihydroindol-2-ones as receptor tyrosine kinase inhibitors.

INVENTOR(S): McNutt, Robert Walton, Jr.; Jung, David Kendall;  
 Harris, Philip Anthony; Hunter, Robert Neil, III;  
 Veal, James Marvin; Dickerson, Scott; Lackey, Karen Elizabeth; Peel, Michael Robert

PATENT ASSIGNEE(S): Glaxo Group Limited, UK

SOURCE: PCT Int. Appl., 144 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

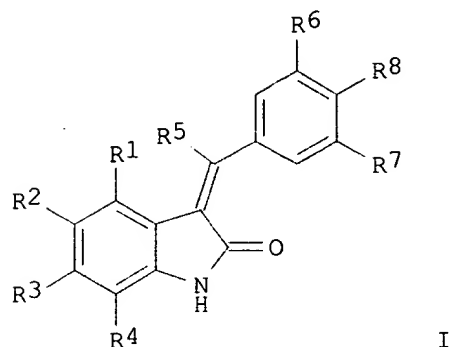
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9910325	A1	19990304	WO 1998-EP4844	19980804
W:	AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,			



FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,  
 CM, GA, GN, GW, ML, MR, NE, SN, TD, TG  
 AU 9891584 A1 19990316 AU 1998-91584 19980804  
 EP 1003721 A1 20000531 EP 1998-943832 19980804  
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,  
 IE, FI  
 US 6268391 B1 20010731 US 2000-446586 20000407  
 PRIORITY APPLN. INFO.: GB 1997-16557 A 19970806  
 WO 1998-EP4844 W 19980804  
 OTHER SOURCE(S): MARPAT 130:209599  
 GI



AB Title compds. [I; R1 = H; R1R2 = fused 5-10 membered aryl, heteroaryl, heterocyclyl; R2, R3 = H, HET, aryl, alipharyl, cyano, NO<sub>2</sub>, halo, R10, OR10, SR10, SOR10, SO<sub>2</sub>R10, NR10R11, etc.; R4 = H, halo, NO<sub>2</sub>, cyano; R5 = H, (substituted) alipharyl; R6, R7 = halo, cyano, NO<sub>2</sub>, CONR10R11, SO<sub>2</sub>NR10R11, NR10R11, OR11; R8 = OH, NHSO<sub>2</sub>R12, NHCOCF<sub>3</sub>; R10 = H, halo, (substituted) alipharyl, aryl, HET; R11 = H, R10; R12 = H, (substituted) alipharyl, HET; HET = benzofuryl, benzoxazolyl, dioxanyl, dithianyl, dithiazinyl, furyl, imidazolyl, indolyl, indazolyl, morpholinyl, tetrazolyl, pyrrolyl, quinolinyl, triazinyl, tetrahydrofuryl, etc.], were prepd. for treatment of tumor growth, preventing organ transplant rejection, healing chronic wounds, etc. (no data). Thus, 5-(2-methylthiazol-4-yl)-1,3-dihydroindol-2-one hydrochloride (prepn. given) was stirred with 3,5-dibromo-4-hydroxybenzaldehyde in AcOH/aq. HCl to give 64% 3-(3,5-dibromo-4-hydroxybenzylidene)-5-(2-methylthiazol-4-yl)-1,3-dihydroindol-2-one.

IT 220904-54-1P 220904-55-2P 220904-56-3P  
 220904-57-4P 220904-58-5P 220904-59-6P  
 220904-60-9P 220904-61-0P 220904-62-1P  
 220904-63-2P 220904-64-3P 220904-65-4P  
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 220905-06-6P

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (prepn. of benzylidene-1,3-dihydroindol-2-ones as receptor tyrosine kinase inhibitors.)

IT 220904-99-4

RL: RCT (Reactant)  
 (prepn. of benzylidene-1,3-dihydroindol-2-ones as receptor tyrosine kinase inhibitors.)

REFERENCE COUNT: 4

REFERENCE(S): (1) Merck & Co Inc; GB 2306108 A 1997 HCAPLUS  
 (2) Sugen Inc; WO 9640116 A 1996 HCAPLUS  
 (3) Sugen Inc; WO 9807695 A 1998 HCAPLUS  
 (4) Tatho Pharmaceutical Co Ltd; WO 9725986 A 1997 HCAPLUS

L18 ANSWER 12 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1998:429042 HCAPLUS

DOCUMENT NUMBER: 129:117426

TITLE: Synthesis and Biological Evaluations of 3-Substituted Indolin-2-ones: A Novel Class of Tyrosine Kinase Inhibitors That Exhibit Selectivity toward Particular Receptor Tyrosine Kinases

AUTHOR(S): Sun, Li; Tran, Ngoc; Tang, Flora; App, Harald; Hirth, Peter; McMahon, Gerald; Tang, Cho

CORPORATE SOURCE: SUGEN Inc, Redwood City, CA, 94063, USA

SOURCE: J. Med. Chem. (1998), 41(14), 2588-2603

CODEN: JMCMAR; ISSN: 0022-2623

PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

AB 3-Substituted indolin-2-ones have been designed and synthesized as a novel class of tyrosine kinase inhibitors which exhibit selectivity toward different receptor tyrosine kinases (RTKs). These compds. have been evaluated for their relative inhibitory properties against a panel of RTKs in intact cells. By modifying the 3-substituted indolin-2-ones, we have identified compds. which showed selective inhibition of the ligand-dependent autophosphorylation of various RTKs at submicromolar levels in cells. Structure-activity anal. for these compds. and their relative potency and selectivity to inhibit particular RTKs has detd. that (1) 3-[(five-membered heteroaryl ring)methylidenyl]indolin-2-ones are highly specific against the VEGF (Flk-1) RTK activity, (2) 3-(substituted benzylidenyl)indolin-2-ones contg. bulky group(s) in the Ph ring at the C-3 position of indolin-2-ones showed high selectivity toward the EGF and Her-2 RTKs, and (3) the compd. contg. an extended side chain at the C-3 position of the indolin-2-one exhibited high potency and selectivity when tested against the PDGF and VEGF (Flk-1) RTKs. Recent published crystallog. data for two of these 3-substituted indolin-2-ones provides a rationale to suggest that these compds. may bind in the ATP binding pocket of RTKs. The structure-activity anal. supports the use of subsets of these compds. as specific chem. leads for the development of RTK-specific drugs with broad application for the treatment of human diseases.

IT 29551-48-2P 40811-69-6P 64264-45-5P  
 64264-56-8P 90828-11-8P 90828-16-3P  
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 210303-16-5P 210303-18-7P 210303-20-1P  
 210303-22-3P 210303-24-5P 210303-26-7P  
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 210303-66-5P 210303-67-6P 210303-78-9P

RL: BAC (Biological activity or effector, except adverse); PRP  
 (Properties); SPN (Synthetic preparation); BIOL (Biological study); PREP  
 (Preparation)

(prepn. and evaluation of 3-substituted indolin-2-ones as  
 inhibitors of selective growth factor receptors)

L18 ANSWER 13 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1998:147306 HCAPLUS

DOCUMENT NUMBER: 128:204803

TITLE: Indolinone combinatorial libraries and related  
 products and methods for the treatment of disease  
 INVENTOR(S): Tang, Peng Cho; Sun, Li; McMahon, Gerald; Hirth, Klaus  
 Peter; Shawver, Laura Kay; et al.

PATENT ASSIGNEE(S): Sugan, Inc., USA; Tang, Peng Cho; Sun, Li; McMahon,  
 Gerald

SOURCE: PCT Int. Appl., 293 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

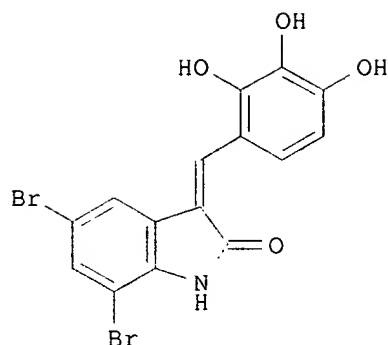
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9807695	A1	19980226	WO 1997-US14736	19970820
W:	AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG			
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R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI			
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JP 2001503736	T2	20010321	JP 1998-510973	19970820
AU 9741556	A1	19980306	AU 1997-41556	19970821
PRIORITY APPLN. INFO.:			US 1996-702232	A 19960823
			US 1996-31585	P 19961205
			US 1996-31586	P 19961205
			US 1996-31588	P 19961205
			US 1996-32546	P 19961205
			US 1996-32547	P 19961205
			US 1997-45565	P 19970505
			US 1997-45566	P 19970505

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US 1997-45715	P	19970505
US 1997-46843	P	19970505
US 1996-45715	P	19961205
US 1997-31565	P	19970505
WO 1997-US14736	W	19970820

OTHER SOURCE(S): MARPAT 128:204803  
GI



AB The invention relates to indolinone derivs. capable of modulating, regulating, and/or inhibiting protein kinase signal transduction. The compds. are useful for the treatment of diseases related to unregulated protein kinase signal transduction, including cell proliferative diseases such as cancer, atherosclerosis, arthritis, and restenosis, and metabolic diseases such as diabetes. Inhibitors specific to the **FLK** protein kinase can be obtained by adding chem. substituents to the 3-[(indole-3-yl)methylene]-2-indolinone system, in particular at the 1' position of the indole ring. Indolinone compds. that specifically inhibit the **FLK** and platelet derived growth factor protein kinases can harbor a tetrahydroindole or cyclopentano[b]pyrrole moiety. Indolinone compds. that are modified with substituents, particularly at the 5 position of the oxindole ring, can effectively activate protein kinases. This invention also features novel hydrosol. indolinone compds. that are tyrosine kinase inhibitors, and related products and methods. Approx. 1200 title compds., such as I, were prepd. by combinatorial condensation of certain (un)substituted indolinones with aldehydes at the 3-position. I gave complete inhibition of MET kinase at chimeric MET receptors in vitro.

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 203988-86-7P,

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 3-[[2-(Ethoxycarbonyl)-3-[2-(ethoxycarbonyl)ethyl]-4-  
 [(ethoxycarbonyl)methyl]pyrrol-5-yl)methylidenyl]-5-bromo-4-methyl-2-

indolinone 203989-28-0P, 3-[[2-Methyl-3-(ethoxycarbonyl)furan-5-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203989-29-1P, 3-[[2,3-Bis(methoxycarbonyl)-5-methylpyrrol-4-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203989-30-4P, 3-(4-Chloro-3-nitrobenzylidenyl)-5-bromo-4-methyl-2-indolinone 203989-31-5P, 3-(2,4-Dihydroxy-3-methylbenzylidenyl)-5-bromo-4-methyl-2-indolinone 203989-32-6P, 3-(Furan-2-ylmethylidenyl)-5-bromo-4-methyl-2-indolinone 203989-33-7P, 3-[(2-Nitrofuran-5-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203989-35-9P, 3-[[2,4-Dimethyl-3-(ethoxycarbonyl)pyrrol-5-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203989-36-0P, 3-(3-Bromo-2-hydroxy-5-methoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203989-38-2P, 3-[(1-Hydroxynaphth-2-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203989-40-6P, 3-[[2-(Ethoxycarbonyl)-3-[2-(ethoxycarbonyl)ethyl]-4-[(ethoxycarbonyl)methyl]pyrrol-5-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203989-42-8P, 3-[[2-Methyl-3-(ethoxycarbonyl)furan-5-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203989-43-9P, 3-[[2,3-Bis(methoxycarbonyl)-5-methylpyrrol-4-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203989-45-1P, 3-(4-Chloro-3-nitrobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203989-46-2P, 3-(2,4-Dihydroxy-3-methylbenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203989-48-4P, 3-(Furan-2-ylmethylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203989-49-5P, 3-[(2-Nitrofuran-5-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203989-52-0P, 3-[[2,4-Dimethyl-3-(ethoxycarbonyl)pyrrol-5-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203989-53-1P, 3-(3-Bromo-2-hydroxy-5-methoxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203989-55-3P, 3-[(1-Hydroxynaphth-2-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203989-56-4P, 3-[[2-(Ethoxycarbonyl)-3-[2-(ethoxycarbonyl)ethyl]-4-[(ethoxycarbonyl)methyl]pyrrol-5-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203989-57-5P, 3-[[2-Methyl-3-(ethoxycarbonyl)furan-5-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203989-58-6P, 3-[[2,3-Bis(methoxycarbonyl)-5-methylpyrrol-4-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203989-60-0P, 3-(4-Chloro-3-nitrobenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203989-61-1P, 3-(2,4-Dihydroxy-3-methylbenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203989-62-2P, 3-(Furan-2-ylmethylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203989-64-4P, 3-[(2-Nitrofuran-5-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203989-65-5P, 3-[[2,4-Dimethyl-3-(ethoxycarbonyl)pyrrol-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203989-66-6P, 3-(3-Bromo-2-hydroxy-5-methoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203989-67-7P, 3-[(1-Hydroxynaphth-2-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203989-68-8P, 3-[[2-(Ethoxycarbonyl)-3-[2-(ethoxycarbonyl)ethyl]-4-[(ethoxycarbonyl)methyl]pyrrol-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203989-69-9P, 3-[[2-Methyl-3-(ethoxycarbonyl)furan-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203989-70-2P, 3-[[2,3-Bis(methoxycarbonyl)-5-methylpyrrol-4-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203989-71-3P, 3-(4-Chloro-3-nitrobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone

203989-72-4P, 3-(2,4-Dihydroxy-3-methylbenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203989-73-5P,  
 3-(Furan-2-ylmethylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203989-74-6P, 3-[(2-Nitrofuran-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203989-75-7P,  
 3-[[2,4-Dimethyl-3-(ethoxycarbonyl)pyrrol-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203989-76-8P, 3-(3-Bromo-2-hydroxy-5-methoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203989-77-9P,  
 3-[(1-Hydroxynaphth-2-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203989-78-0P, 3-[[2-(Ethoxycarbonyl)-3-[2-(ethoxycarbonyl)ethyl]-4-[(ethoxycarbonyl)methyl]pyrrol-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203989-79-1P, 3-[[2-Methyl-3-(ethoxycarbonyl)furan-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203989-80-4P,  
 3-[[2,3-Bis(methoxycarbonyl)-5-methylpyrrol-4-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203989-81-5P, 3-(4-Chloro-3-nitrobenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203989-82-6P,  
 3-(2,4-Dihydroxy-3-methylbenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203989-83-7P, 3-(Furan-2-ylmethylidenyl)-5-(2-chloroethyl)-2-indolinone 203989-84-8P, 3-[(2-Nitrofuran-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203989-85-9P, 3-(4-Ethoxy-3-methoxybenzylidenyl)-5,7-dibromo-2-indolinone 203989-86-0P,  
 3-(3,4-Dihydroxybenzylidenyl)-5,7-dibromo-2-indolinone 203989-87-1P, 3-(2,4-Dimethoxybenzylidenyl)-5,7-dibromo-2-indolinone 203989-88-2P, 3-[(2,4-Dimethyl-3-ethylpyrrol-5-yl)methylidenyl]-5,7-dibromo-2-indolinone 203989-89-3P,  
 3-(2,4,6-Trimethoxybenzylidenyl)-5,7-dibromo-2-indolinone 203989-90-6P, 3-(4-Hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 203989-91-7P, 3-[4-(Dimethylamino)benzylidenyl]-5,7-dibromo-2-indolinone 203989-92-8P, 3-(2-Chloro-4-fluorobenzylidenyl)-5,7-dibromo-2-indolinone 203989-93-9P, 3-(3-Nitrobenzylidenyl)-5,7-dibromo-2-indolinone 203989-94-0P, 3-[4-Fluoro-2-(trifluoromethyl)benzylidenyl]-5,7-dibromo-2-indolinone 203989-95-1P,  
 3-(4-Ethoxy-3-methoxybenzylidenyl)-5-iodo-2-indolinone 203989-96-2P, 3-(3,4-Dihydroxybenzylidenyl)-5-iodo-2-indolinone 203989-97-3P, 3-(2,4-Dimethoxybenzylidenyl)-5-iodo-2-indolinone 203989-98-4P, 3-[(2,4-Dimethyl-3-ethylpyrrol-5-yl)methylidenyl]-5-iodo-2-indolinone 203989-99-5P,  
 3-(2,4,6-Trimethoxybenzylidenyl)-5-iodo-2-indolinone 203990-00-5P, 3-(4-Hydroxybenzylidenyl)-5-iodo-2-indolinone 203990-01-6P,  
 3-[4-(Dimethylamino)benzylidenyl]-5-iodo-2-indolinone 203990-02-7P, 3-(2-Chloro-4-fluorobenzylidenyl)-5-iodo-2-indolinone 203990-03-8P, 3-(3-Nitrobenzylidenyl)-5-iodo-2-indolinone 203990-04-9P, 3-[4-Fluoro-2-(trifluoromethyl)benzylidenyl]-5-iodo-2-indolinone 203990-05-0P, 3-(4-Ethoxy-3-methoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-06-1P, 3-(3,4-Dihydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-07-2P,  
 3-(2,4-Dimethoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-08-3P, 3-[(2,4-Dimethyl-3-ethylpyrrol-5-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203990-09-4P, 3-(2,4,6-Trimethoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-10-7P,  
 3-(4-Hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-11-8P, 3-[4-(Dimethylamino)benzylidenyl]-5-bromo-4-methyl-2-indolinone 203990-12-9P, 3-(2-Chloro-4-fluorobenzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-13-0P, 3-(3-Nitrobenzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-14-1P,  
 3-[4-Fluoro-2-(trifluoromethyl)benzylidenyl]-5-bromo-4-methyl-2-indolinone 203990-15-2P, 3-(4-Ethoxy-3-methoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203990-16-3P,  
 3-(3,4-Dihydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone



203990-17-4P, 3-(2,4-Dimethoxybenzylidenyl)-5-  
 [(methylamino)sulfonyl]-2-indolinone 203990-18-5P,  
 3-[(2,4-Dimethyl-3-ethylpyrrol-5-yl)methylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203990-19-6P,  
 3-(2,4,6-Trimethoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone  
 203990-20-9P, 3-(4-Hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-  
 2-indolinone 203990-21-0P, 3-[4-(Dimethylamino)benzylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203990-22-1P,  
 3-(2-Chloro-4-fluorobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone  
 203990-23-2P, 3-(3-Nitrobenzylidenyl)-5-[(methylamino)sulfonyl]-2-  
 indolinone 203990-24-3P, 3-[4-Fluoro-2-  
 (trifluoromethyl)benzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone  
 203990-25-4P, 3-(4-Ethoxy-3-methoxybenzylidenyl)-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203990-26-5P  
 , 3-(3,4-Dihydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfo  
 nyl]-2-indolinone 203990-27-6P, 3-(2,4-Dimethoxybenzylidenyl)-5-  
 [[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone  
 203990-28-7P, 3-[(2,4-Dimethyl-3-ethylpyrrol-5-yl)methylidenyl]-5-  
 [[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone  
 203990-29-8P, 3-(2,4,6-Trimethoxybenzylidenyl)-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203990-30-1P  
 , 3-(4-Hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-  
 2-indolinone 203990-31-2P, 3-[4-(Dimethylamino)benzylidenyl]-5-  
 [[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone  
 203990-32-3P  
 , 3-(2-Chloro-4-fluorobenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfo  
 nyl]-2-indolinone 203990-33-4P, 3-(3-Nitrobenzylidenyl)-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203990-34-5P  
 , 3-[4-Fluoro-2-(trifluoromethyl)benzylidenyl]-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203990-35-6P  
 , 3-(4-Ethoxy-3-methoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone  
 203990-36-7P, 3-(3,4-Dihydroxybenzylidenyl)-5-(morpholinosulfonyl)-  
 2-indolinone 203990-37-8P, 3-(2,4-Dimethoxybenzylidenyl)-5-  
 (morpholinosulfonyl)-2-indolinone 203990-38-9P,  
 3-[(2,4-Dimethyl-3-ethylpyrrol-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-  
 indolinone 203990-39-0P, 3-(2,4,6-Trimethoxybenzylidenyl)-5-  
 (morpholinosulfonyl)-2-indolinone 203990-40-3P,  
 3-(4-Hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone  
 203990-41-4P, 3-[4-(Dimethylamino)benzylidenyl]-5-  
 (morpholinosulfonyl)-2-indolinone 203990-42-5P,  
 3-(2-Chloro-4-fluorobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone  
 203990-43-6P, 3-(3-Nitrobenzylidenyl)-5-(morpholinosulfonyl)-2-  
 indolinone 203990-44-7P, 3-[4-Fluoro-2-  
 (trifluoromethyl)benzylidenyl]-5-(morpholinosulfonyl)-2-indolinone  
 203990-45-8P, 3-(4-Ethoxy-3-methoxybenzylidenyl)-5-(2-chloroethyl)-  
 2-indolinone 203990-46-9P, 3-(3,4-Dihydroxybenzylidenyl)-5-(2-  
 chloroethyl)-2-indolinone 203990-47-0P, 3-(2,4-  
 Dimethoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203990-48-1P  
 , 3-[(2,4-Dimethyl-3-ethylpyrrol-5-yl)methylidenyl]-5-(2-chloroethyl)-2-  
 indolinone 203990-49-2P, 3-(2,4,6-Trimethoxybenzylidenyl)-5-(2-  
 chloroethyl)-2-indolinone 203990-50-5P, 3-(4-  
 Hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203990-51-6P  
 , 3-[4-(Dimethylamino)benzylidenyl]-5-(2-chloroethyl)-2-indolinone  
 203990-52-7P, 3-(2-Chloro-4-fluorobenzylidenyl)-5-(2-chloroethyl)-  
 2-indolinone 203990-53-8P, 3-(3-Nitrobenzylidenyl)-5-(2-  
 chloroethyl)-2-indolinone 203990-54-9P, 3-[4-Fluoro-2-  
 (trifluoromethyl)benzylidenyl]-5-(2-chloroethyl)-2-indolinone  
 203990-55-0P, 3-(2,4,6-Trifluorobenzylidenyl)-5,7-dibromo-2-  
 indolinone 203990-56-1P, 3-(4-Hydroxy-2-methoxybenzylidenyl)-5,7-

dibromo-2-indolinone 203990-57-2P, 3-(3,4-Dimethoxybenzylidenyl)-5,7-dibromo-2-indolinone 203990-58-3P, 3-(2-Hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 203990-59-4P, 3-(Benzylidenyl)-5,7-dibromo-2-indolinone 203990-60-7P, 3-[[2-(Methylthio)thien-5-yl)methylidenyl]-5,7-dibromo-2-indolinone 203990-61-8P, 3-(2,4-Dihydroxy-6-methylbenzylidenyl)-5,7-dibromo-2-indolinone 203990-62-9P, 3-(3-Ethoxy-4-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 203990-63-0P, 3-(2-Hydroxy-5-methoxybenzylidenyl)-5,7-dibromo-2-indolinone 203990-64-1P, 3-[(Imidazol-2-yl)methylidenyl]-5,7-dibromo-2-indolinone 203990-65-2P, 3-(2,4,6-Trifluorobenzylidenyl)-5-iodo-2-indolinone 203990-66-3P, 3-(4-Hydroxy-2-methoxybenzylidenyl)-5-iodo-2-indolinone 203990-67-4P, 3-(3,4-Dimethoxybenzylidenyl)-5-iodo-2-indolinone 203990-68-5P, 3-(2-Hydroxybenzylidenyl)-5-iodo-2-indolinone 203990-69-6P, 3-(Benzylidenyl)-5-iodo-2-indolinone 203990-70-9P, 3-[[2-(Methylthio)thien-5-yl)methylidenyl]-5-iodo-2-indolinone 203990-71-0P, 3-(2,4-Dihydroxy-6-methylbenzylidenyl)-5-iodo-2-indolinone 203990-72-1P, 3-(3-Ethoxy-4-hydroxybenzylidenyl)-5-iodo-2-indolinone 203990-73-2P, 3-(2-Hydroxy-5-methoxybenzylidenyl)-5-iodo-2-indolinone  
 RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. and testing of indolinone combinatorial library as protein kinase inhibitors)

IT 203990-74-3P, 3-[(Imidazol-2-yl)methylidenyl]-5-iodo-2-indolinone  
 203990-75-4P, 3-(2,4,6-Trifluorobenzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-76-5P, 3-(4-Hydroxy-2-methoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-77-6P, 3-(3,4-Dimethoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-78-7P, 3-(2-Hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-79-8P, 3-(Benzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-80-1P, 3-[[2-(Methylthio)thien-5-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203990-81-2P, 3-(2,4-Dihydroxy-6-methylbenzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-82-3P, 3-(3-Ethoxy-4-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-83-4P, 3-(2-Hydroxy-5-methoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203990-84-5P, 3-[(Imidazol-2-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203990-85-6P, 3-(2,4,6-Trifluorobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203990-86-7P, 3-(4-Hydroxy-2-methoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203990-87-8P, 3-(3,4-Dimethoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203990-88-9P, 3-(2-Hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203990-89-0P, 3-(Benzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203990-90-3P, 3-[[2-(Methylthio)thien-5-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203990-91-4P, 3-(2,4-Dihydroxy-6-methylbenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203990-92-5P, 3-(3-Ethoxy-4-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203990-93-6P, 3-(2-Hydroxy-5-methoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203990-94-7P, 3-[(Imidazol-2-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203990-96-9P, 3-(2,4,6-Trifluorobenzylidenyl)-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203990-98-1P, 3-(4-Hydroxy-2-methoxybenzylidenyl)-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203991-00-8P, 3-(3,4-Dimethoxybenzylidenyl)-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203991-02-0P, 3-(2-Hydroxybenzylidenyl)-5-[[4-

(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203991-04-2P,  
 3-(Benzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203991-05-3P, 3-[[2-(Methylthio)thien-5-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203991-07-5P, 3-(2,4-Dihydroxy-6-methylbenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203991-10-0P, 3-(3-Ethoxy-4-hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203991-13-3P, 3-(2-Hydroxy-5-methoxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203991-16-6P, 3-[(Imidazol-2-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203991-20-2P, 3-(2,4,6-Trifluorobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203991-23-5P, 3-(4-Hydroxy-2-methoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203991-27-9P, 3-(3,4-Dimethoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203991-31-5P, 3-(2-Hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203991-32-6P, 3-(Benzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203991-33-7P, 3-[[2-(Methylthio)thien-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203991-34-8P, 3-(2,4-Dihydroxy-6-methylbenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203991-35-9P, 3-(3-Ethoxy-4-hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203991-37-1P, 3-(2-Hydroxy-5-methoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203991-39-3P, 3-[(Imidazol-2-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203991-41-7P, 3-(2,4,6-Trifluorobenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203991-43-9P, 3-(4-Hydroxy-2-methoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203991-45-1P, 3-(3,4-Dimethoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203991-47-3P, 3-(2-Hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203991-49-5P, 3-(Benzylidenyl)-5-(2-chloroethyl)-2-indolinone 203991-51-9P, 3-[[2-(Methylthio)thien-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203991-52-0P, 3-(2,4-Dihydroxy-6-methylbenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203991-53-1P, 3-(3-Ethoxy-4-hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203991-54-2P, 3-(2-Hydroxy-5-methoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203991-55-3P, 3-[(Imidazol-2-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203991-56-4P, 3-[(1-Methylbenzimidazol-2-yl)methylidenyl]-5,7-dibromo-2-indolinone 203991-57-5P, 3-[(4-Chloro-1-methylpyrazol-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203991-58-6P, 3-[(2,3-Dimethylthien-5-yl)methylidenyl]-5,7-dibromo-2-indolinone 203991-59-7P, 3-[(4,5,6,7-Tetrahydroindol-2-yl)methylidenyl]-5,7-dibromo-2-indolinone 203991-60-0P, 3-[3-(Chloromethyl)-2-hydroxy-5-nitrobenzylidenyl]-5,7-dibromo-2-indolinone 203991-61-1P, 3-[(2-Chlorothien-5-yl)methylidenyl]-5,7-dibromo-2-indolinone 203991-62-2P, 3-[(2,4-Dimethylpyrrol-5-yl)methylidenyl]-5,7-dibromo-2-indolinone 203991-63-3P, 3-(3-tert-Butyl-4-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 203991-64-4P, 3-(3-Bromo-5-tert-butyl-4-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 203991-65-5P, 3-(3,5-Di-tert-butyl-4-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 203991-66-6P, 3-[(1-Methylbenzimidazol-2-yl)methylidenyl]-5-iodo-2-indolinone 203991-67-7P, 3-[(4-Chloro-1-methylpyrazol-3-yl)methylidenyl]-5-iodo-2-indolinone 203991-68-8P, 3-[(2,3-Dimethylthien-5-yl)methylidenyl]-5-iodo-2-indolinone 203991-69-9P, 3-[(4,5,6,7-Tetrahydroindol-2-yl)methylidenyl]-5-iodo-2-indolinone 203991-70-2P, 3-[3-(Chloromethyl)-2-hydroxy-5-nitrobenzylidenyl]-5-iodo-2-indolinone 203991-71-3P, 3-[(2-Chlorothien-5-yl)methylidenyl]-5-iodo-2-indolinone

203991-72-4P, 3-[(2,4-Dimethylpyrrol-5-yl)methylidenyl]-5-iodo-2-indolinone 203991-73-5P, 3-(3-tert-Butyl-4-hydroxybenzylidenyl)-5-iodo-2-indolinone 203991-74-6P, 3-(3-Bromo-5-tert-butyl-4-hydroxybenzylidenyl)-5-iodo-2-indolinone 203991-75-7P, 3-(3,5-Di-tert-butyl-4-hydroxybenzylidenyl)-5-iodo-2-indolinone 203991-76-8P, 3-[(1-Methylbenzimidazol-2-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203991-77-9P, 3-[(4-Chloro-1-methylpyrazol-3-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203991-78-0P, 3-[(2,3-Dimethylthien-5-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203991-79-1P, 3-[(4,5,6,7-Tetrahydroindol-2-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203991-80-4P, 3-[3-(Chloromethyl)-2-hydroxy-5-nitrobenzylidenyl]-5-bromo-4-methyl-2-indolinone 203991-81-5P, 3-[(2-Chlorothien-5-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203991-82-6P, 3-[(2,4-Dimethylpyrrol-5-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203991-83-7P, 3-(3-tert-Butyl-4-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203991-84-8P, 3-(3-Bromo-5-tert-butyl-4-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203991-85-9P, 3-(3,5-Di-tert-butyl-4-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203991-86-0P, 3-[(1-Methylbenzimidazol-2-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203991-87-1P, 3-[(4-Chloro-1-methylpyrazol-3-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203991-88-2P, 3-[(2,3-Dimethylthien-5-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203991-89-3P, 3-[(4,5,6,7-Tetrahydroindol-2-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203991-90-6P, 3-[3-(Chloromethyl)-2-hydroxy-5-nitrobenzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203991-91-7P, 3-[(2-Chlorothien-5-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203991-92-8P, 3-[(2,4-Dimethylpyrrol-5-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203991-93-9P, 3-(3-tert-Butyl-4-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203991-94-0P, 3-(3-Bromo-5-tert-butyl-4-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203991-95-1P, 3-(3,5-Di-tert-butyl-4-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203991-96-2P, 3-[(1-Methylbenzimidazol-2-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203991-97-3P, 3-[(4-Chloro-1-methylpyrazol-3-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203991-98-4P, 3-[(2,3-Dimethylthien-5-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203991-99-5P, 3-[(4,5,6,7-Tetrahydroindol-2-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-00-1P, 3-[3-(Chloromethyl)-2-hydroxy-5-nitrobenzylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-01-2P, 3-[(2-Chlorothien-5-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-02-3P, 3-[(2,4-Dimethylpyrrol-5-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-03-4P, 3-(3-tert-Butyl-4-hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-04-5P, 3-(3-Bromo-5-tert-butyl-4-hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-05-6P, 3-(3,5-Di-tert-butyl-4-hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-06-7P, 3-[(1-Methylbenzimidazol-2-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203992-07-8P, 3-[(4-Chloro-1-methylpyrazol-3-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203992-08-9P, 3-[(2,3-Dimethylthien-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-

indolinone 203992-09-0P, 3-[(4,5,6,7-Tetrahydroindol-2-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203992-10-3P,  
3-[3-(Chloromethyl)-2-hydroxy-5-nitrobenzylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203992-11-4P,  
3-[(2-Chlorothien-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203992-12-5P, 3-[(2,4-Dimethylpyrrol-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203992-13-6P,  
3-(3-tert-Butyl-4-hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203992-14-7P, 3-(3-Bromo-5-tert-butyl-4-hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203992-15-8P,  
3-(3,5-Di-tert-butyl-4-hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203992-16-9P, 3-[(1-Methylbenzimidazol-2-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203992-17-0P,  
3-[(4-Chloro-1-methylpyrazol-3-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203992-18-1P, 3-[(2,3-Dimethylthien-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203992-19-2P,  
3-[(4,5,6,7-Tetrahydroindol-2-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203992-20-5P, 3-[3-(Chloromethyl)-2-hydroxy-5-nitrobenzylidenyl]-5-(2-chloroethyl)-2-indolinone 203992-21-6P,  
3-[(2-Chlorothien-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203992-22-7P, 3-[(2,4-Dimethylpyrrol-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203992-23-8P, 3-(3-tert-Butyl-4-hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203992-24-9P,  
3-(3-Bromo-5-tert-butyl-4-hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203992-25-0P, 3-(3,5-Di-tert-butyl-4-hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203992-26-1P,  
3-(3-tert-Butyl-4-hydroxy-5-nitrobenzylidenyl)-5,7-dibromo-2-indolinone 203992-27-2P, 3-(2,4,6-Trihydroxybenzylidenyl)-5,7-dibromo-2-indolinone 203992-28-3P, 3-[(2-Nitrothien-5-yl)methylidenyl]-5,7-dibromo-2-indolinone 203992-29-4P, 3-(4-Carboxybenzylidenyl)-5,7-dibromo-2-indolinone 203992-30-7P, 3-(2,4-Difluorobenzylidenyl)-5,7-dibromo-2-indolinone 203992-31-8P, 3-(3,5-Dimethyl-4-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 203992-32-9P,  
3-(3-tert-Butyl-5-chloro-4-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 203992-33-0P, 3-[(2-Nitrothien-4-yl)methylidenyl]-5,7-dibromo-2-indolinone 203992-34-1P, 3-[4-(Di-n-butylamino)benzylidenyl]-5,7-dibromo-2-indolinone 203992-35-2P, 3-[4-(Trifluoromethyl)benzylidenyl]-5,7-dibromo-2-indolinone 203992-36-3P, 3-(3-tert-Butyl-4-hydroxy-5-nitrobenzylidenyl)-5-iodo-2-indolinone 203992-37-4P, 3-(2,4,6-Trihydroxybenzylidenyl)-5-iodo-2-indolinone 203992-38-5P, 3-[(2-Nitrothien-5-yl)methylidenyl]-5-iodo-2-indolinone 203992-39-6P,  
3-(4-Carboxybenzylidenyl)-5-iodo-2-indolinone 203992-40-9P, 3-(2,4-Difluorobenzylidenyl)-5-iodo-2-indolinone 203992-41-0P, 3-(3,5-Dimethyl-4-hydroxybenzylidenyl)-5-iodo-2-indolinone 203992-42-1P, 3-(3-tert-Butyl-5-chloro-4-hydroxybenzylidenyl)-5-iodo-2-indolinone 203992-43-2P, 3-[(2-Nitrothien-4-yl)methylidenyl]-5-iodo-2-indolinone 203992-44-3P,  
3-[4-(Di-n-butylamino)benzylidenyl]-5-iodo-2-indolinone 203992-45-4P, 3-[4-(Trifluoromethyl)benzylidenyl]-5-iodo-2-indolinone 203992-46-5P, 3-(3-tert-Butyl-4-hydroxy-5-nitrobenzylidenyl)-5-bromo-4-methyl-2-indolinone 203992-47-6P,  
3-(2,4,6-Trihydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203992-48-7P, 3-[(2-Nitrothien-5-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203992-49-8P, 3-(4-Carboxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203992-50-1P, 3-(2,4-Difluorobenzylidenyl)-5-bromo-4-methyl-2-indolinone 203992-51-2P, 3-(3,5-Dimethyl-4-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203992-52-3P,  
3-(3-tert-Butyl-5-chloro-4-hydroxybenzylidenyl)-5-bromo-4-methyl-2-

indolinone 203992-53-4P, 3-[(2-Nitrothien-4-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203992-54-5P, 3-[4-(Di-n-butylamino)benzylidenyl]-5-bromo-4-methyl-2-indolinone 203992-55-6P, 3-[4-(Trifluoromethyl)benzylidenyl]-5-bromo-4-methyl-2-indolinone 203992-56-7P, 3-(3-tert-Butyl-4-hydroxy-5-nitrobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203992-57-8P, 3-(2,4,6-Trihydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203992-58-9P, 3-[(2-Nitrothien-5-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203992-59-0P, 3-(4-Carboxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203992-60-3P, 3-(2,4-Difluorobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203992-61-4P, 3-(3,5-Dimethyl-4-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203992-62-5P, 3-(3-tert-Butyl-5-chloro-4-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203992-63-6P, 3-[(2-Nitrothien-4-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203992-64-7P, 3-[4-(Di-n-butylamino)benzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203992-65-8P, 3-[4-(Trifluoromethyl)benzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203992-66-9P, 3-(3-tert-Butyl-4-hydroxy-5-nitrobenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-67-0P\*\*\*, 3-(2,4,6-Trihydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone \*\*\*203992-68-1P, 3-[(2-Nitrothien-5-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-69-2P, 3-(4-Carboxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-70-5P, 3-(2,4-Difluorobenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-71-6P, 3-(3,5-Dimethyl-4-hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-72-7P, 3-(3-tert-Butyl-5-chloro-4-hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-73-8P, 3-[(2-Nitrothien-4-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-74-9P, 3-[4-(Di-n-butylamino)benzylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-75-0P, 3-[4-(Trifluoromethyl)benzylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203992-76-1P, 3-(3-tert-Butyl-4-hydroxy-5-nitrobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203992-77-2P, 3-(2,4,6-Trihydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203992-78-3P, 3-[(2-Nitrothien-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203992-79-4P, 3-(4-Carboxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203992-80-7P, 3-(2,4-Difluorobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203992-81-8P, 3-(3,5-Dimethyl-4-hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203992-82-9P, 3-(3-tert-Butyl-5-chloro-4-hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203992-83-0P, 3-[(2-Nitrothien-4-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203992-84-1P, 3-[4-(Di-n-butylamino)benzylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203992-85-2P, 3-[4-(Trifluoromethyl)benzylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203992-86-3P, 3-(3-tert-Butyl-4-hydroxy-5-nitrobenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203992-87-4P, 3-(2,4,6-Trihydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203992-88-5P, 3-[(2-Nitrothien-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203992-89-6P, 3-(4-Carboxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203992-90-9P, 3-(2,4-Difluorobenzylidenyl)-5-(2-chloroethyl)-2-indolinone

203992-91-0P, 3-(3,5-Dimethyl-4-hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203992-92-1P, 3-(3-tert-Butyl-5-chloro-4-hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203992-93-2P, 3-[(2-Nitrothien-4-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203992-94-3P, 3-[4-(Di-n-butylamino)benzylidenyl]-5-(2-chloroethyl)-2-indolinone 203992-95-4P, 3-[4-(Trifluoromethyl)benzylidenyl]-5-(2-chloroethyl)-2-indolinone 203992-96-5P, 3-(2,3,4-Trihydroxybenzylidenyl)-5,7-dibromo-2-indolinone 203992-97-6P, 3-(2-Hydroxy-3-methoxybenzylidenyl)-5,7-dibromo-2-indolinone 203992-98-7P, 3-(3-Bromo-4,5-dihydroxybenzylidenyl)-5,7-dibromo-2-indolinone 203992-99-8P, 3-(3,4-Diacetoxybenzylidenyl)-5,7-dibromo-2-indolinone 203993-00-4P, 3-(4-Hydroxy-3-methylbenzylidenyl)-5,7-dibromo-2-indolinone 203993-01-5P, 3-(2-Bromobenzylidenyl)-5,7-dibromo-2-indolinone 203993-02-6P, 3-(2,4-Dihydroxybenzylidenyl)-5,7-dibromo-2-indolinone 203993-03-7P, 3-(2-Hydroxy-4-methoxybenzylidenyl)-5,7-dibromo-2-indolinone 203993-04-8P, 3-(3-Bromobenzylidenyl)-5,7-dibromo-2-indolinone 203993-05-9P, 3-(3,5-Di-tert-butyl-2-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 203993-06-0P, 3-(2,3,4-Trihydroxybenzylidenyl)-5-iodo-2-indolinone 203993-07-1P, 3-(2-Hydroxy-3-methoxybenzylidenyl)-5-iodo-2-indolinone 203993-08-2P, 3-(3-Bromo-4,5-dihydroxybenzylidenyl)-5-iodo-2-indolinone 203993-09-3P, 3-(3,4-Diacetoxybenzylidenyl)-5-iodo-2-indolinone 203993-10-6P, 3-(4-Hydroxy-3-methylbenzylidenyl)-5-iodo-2-indolinone 203993-11-7P, 3-(2-Bromobenzylidenyl)-5-iodo-2-indolinone 203993-12-8P, 3-(2,4-Dihydroxybenzylidenyl)-5-iodo-2-indolinone 203993-13-9P, 3-(2-Hydroxy-4-methoxybenzylidenyl)-5-iodo-2-indolinone 203993-14-0P, 3-(3-Bromobenzylidenyl)-5-iodo-2-indolinone 203993-15-1P, 3-(3,5-Di-tert-butyl-2-hydroxybenzylidenyl)-5-iodo-2-indolinone 203993-16-2P, 3-(2,3,4-Trihydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203993-17-3P, 3-(2-Hydroxy-3-methoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203993-18-4P, 3-(3-Bromo-4,5-dihydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203993-19-5P, 3-(3,4-Diacetoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203993-20-8P, 3-(2-Bromobenzylidenyl)-5-bromo-4-methyl-2-indolinone 203993-21-9P, 3-(2,4-Dihydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203993-22-0P, 3-(2-Hydroxy-4-methoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203993-23-1P, 3-(3-Bromobenzylidenyl)-5-bromo-4-methyl-2-indolinone 203993-24-2P, 3-(3,5-Di-tert-butyl-2-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203993-25-3P, 3-(2,3,4-Trihydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203993-26-4P, 3-(2-Hydroxy-3-methoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203993-27-5P, 3-(3-Bromo-4,5-dihydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203993-28-6P, 3-(3,4-Diacetoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203993-29-7P, 3-(4-Hydroxy-3-methylbenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203993-30-0P, 3-(2-Bromobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203993-31-1P, 3-(2,4-Dihydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203993-32-2P, 3-(2-Hydroxy-4-methoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203993-33-3P, 3-(3-Bromobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203993-34-4P, 3-(3,5-Di-tert-butyl-2-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203993-35-5P, 3-(2,3,4-Trihydroxybenzylidenyl)-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203993-36-6P

, 3-(2-Hydroxy-3-methoxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203993-37-7P**, 3-(3-Bromo-4,5-dihydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203993-38-8P**, 3-(3,4-Diacetoxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203993-39-9P**, 3-(4-Hydroxy-3-methylbenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. and testing of indolinone combinatorial library as protein kinase inhibitors)

IT **203993-40-2P**, 3-(2-Bromobenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203993-41-3P**, 3-(2,4-Dihydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203993-42-4P**, 3-(2-Hydroxy-4-methoxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203993-43-5P**, 3-(3-Bromobenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203993-44-6P**, 3-(3,5-Di-tert-butyl-2-hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203993-45-7P**, 3-(2,3,4-Trihydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203993-46-8P**, 3-(2-Hydroxy-3-methoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203993-47-9P**, 3-(3-Bromo-4,5-dihydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203993-48-0P**, 3-(3,4-Diacetoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203993-49-1P**, 3-(4-Hydroxy-3-methylbenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203993-50-4P**, 3-(2-Bromobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203993-51-5P**, 3-(2,4-Dihydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203993-52-6P**, 3-(2-Hydroxy-4-methoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203993-53-7P**, 3-(3-Bromobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203993-54-8P**, 3-(3,5-Di-tert-butyl-2-hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203993-55-9P**, 3-(2,3,4-Trihydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203993-56-0P**, 3-(2-Hydroxy-3-methoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203993-57-1P**, 3-(3-Bromo-4,5-dihydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203993-58-2P**, 3-(3,4-Diacetoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203993-59-3P**, 3-(4-Hydroxy-3-methylbenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203993-60-6P**, 3-(2-Bromobenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203993-61-7P**, 3-(2,4-Dihydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203993-62-8P**, 3-(2-Hydroxy-4-methoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203993-63-9P**, 3-(3-Bromobenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203993-64-0P**, 3-(3,5-Di-tert-butyl-2-hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203993-65-1P**, 3-[[[1-(Dimethylamino)naphth-4-yl)methylidenyl]-5,7-dibromo-2-indolinone **203993-66-2P**, 3-(4-Hydroxy-3-nitrobenzylidenyl)-5,7-dibromo-2-indolinone **203993-67-3P**, 3-(3-Hydroxy-4-nitrobenzylidenyl)-5,7-dibromo-2-indolinone **203993-68-4P**, 3-[(8-Hydroxy-2,3,6,7-tetrahydro-1H,5H-benzo[ij]quinolizin-9-yl)methylidenyl]-5,7-dibromo-2-indolinone **203993-69-5P**, 3-(3,5-Diisopropyl-4-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone **203993-70-8P**, 3-[(Benzo[b]furan-2-yl)methylidenyl]-5,7-dibromo-2-indolinone **203993-71-9P**, 3-[[[1-(4-Chlorophenyl)pyrrol-2-yl)methylidenyl]-5,7-dibromo-2-indolinone **203993-72-0P**, 3-[(2-Ethylfuran-5-yl)methylidenyl]-5,7-dibromo-2-indolinone **203993-73-1P**,



3-[(3,4-Dimethylthieno[2,3-b]thien-2-yl)methylidenyl]-5,7-dibromo-2-indolinone **203993-74-2P**, 3-[[1-(Dimethylamino)naphth-4-yl)methylidenyl]-5-iodo-2-indolinone **203993-75-3P**, 3-(4-Hydroxy-3-nitrobenzylidenyl)-5-iodo-2-indolinone **203993-76-4P**, 3-(3-Hydroxy-4-nitrobenzylidenyl)-5-iodo-2-indolinone **203993-77-5P**, 3-[(8-Hydroxy-2,3,6,7-tetrahydro-1H,5H-benzo[*ij*]quinolizin-9-yl)methylidenyl]-5-iodo-2-indolinone **203993-78-6P**, 3-(3,5-Diisopropyl-4-hydroxybenzylidenyl)-5-iodo-2-indolinone **203993-79-7P**, 3-[(Benzo[*b*]furan-2-yl)methylidenyl]-5-iodo-2-indolinone **203993-80-0P**, 3-[[1-(4-Chlorophenyl)pyrrol-2-yl)methylidenyl]-5-iodo-2-indolinone **203993-81-1P**, 3-[(2-Ethylfuran-5-yl)methylidenyl]-5-iodo-2-indolinone **203993-82-2P**, 3-[(3,4-Dimethylthieno[2,3-b]thien-2-yl)methylidenyl]-5-iodo-2-indolinone **203993-83-3P**, 3-[[1-(Dimethylamino)naphth-4-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone **203993-84-4P**, 3-(4-Hydroxy-3-nitrobenzylidenyl)-5-bromo-4-methyl-2-indolinone **203993-85-5P**, 3-(3-Hydroxy-4-nitrobenzylidenyl)-5-bromo-4-methyl-2-indolinone **203993-86-6P**, 3-[(8-Hydroxy-2,3,6,7-tetrahydro-1H,5H-benzo[*ij*]quinolizin-9-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone **203993-87-7P**, 3-(3,5-Diisopropyl-4-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone **203993-88-8P**, 3-[(Benzo[*b*]furan-2-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone **203993-89-9P**, 3-[[1-(4-Chlorophenyl)pyrrol-2-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone **203993-90-2P**, 3-[(2-Ethylfuran-5-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone **203993-91-3P**, 3-[(3,4-Dimethylthieno[2,3-b]thien-2-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone **203993-92-4P**, 3-[[1-(Dimethylamino)naphth-4-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone **203993-93-5P**, 3-(4-Hydroxy-3-nitrobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone **203993-94-6P**, 3-(3-Hydroxy-4-nitrobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone **203993-95-7P**, 3-[(8-Hydroxy-2,3,6,7-tetrahydro-1H,5H-benzo[*ij*]quinolizin-9-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone **203993-96-8P**, 3-(3,5-Diisopropyl-4-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone **203993-97-9P**, 3-[(Benzo[*b*]furan-2-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone **203993-98-0P**, 3-[[1-(4-Chlorophenyl)pyrrol-2-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone **203993-99-1P**, 3-[(2-Ethylfuran-5-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone **203994-00-7P**, 3-[(3,4-Dimethylthieno[2,3-b]thien-2-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone **203994-01-8P**, 3-[[1-(Dimethylamino)naphth-4-yl)methylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203994-02-9P**, 3-(4-Hydroxy-3-nitrobenzylidenyl)-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203994-03-0P**, 3-(3-Hydroxy-4-nitrobenzylidenyl)-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203994-04-1P**, 3-[(8-Hydroxy-2,3,6,7-tetrahydro-1H,5H-benzo[*ij*]quinolizin-9-yl)methylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203994-05-2P**, 3-(3,5-Diisopropyl-4-hydroxybenzylidenyl)-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203994-06-3P**, 3-[(Benzo[*b*]furan-2-yl)methylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203994-07-4P**, 3-[[1-(4-Chlorophenyl)pyrrol-2-yl)methylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203994-08-5P**, 3-[(2-Ethylfuran-5-yl)methylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **203994-09-6P**, 3-[(3,4-Dimethylthieno[2,3-b]thien-2-yl)methylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-

2-indolinone 203994-10-9P, 3-[[1-(Dimethylamino)naphth-4-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203994-11-0P,  
 3-(4-Hydroxy-3-nitrobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203994-12-1P, 3-(3-Hydroxy-4-nitrobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203994-13-2P,  
 3-[(8-Hydroxy-2,3,6,7-tetrahydro-1H,5H-benzo[ij]quinolizin-9-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203994-14-3P,  
 3-(3,5-Diisopropyl-4-hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203994-15-4P, 3-[(Benzo[b]furan-2-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203994-16-5P,  
 3-[[1-(4-Chlorophenyl)pyrrol-2-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203994-17-6P, 3-[(2-Ethylfuran-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203994-18-7P,  
 3-[(3,4-Dimethylthieno[2,3-b]thien-2-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203994-19-8P,  
 3-[[1-(Dimethylamino)naphth-4-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203994-20-1P, 3-(4-Hydroxy-3-nitrobenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203994-21-2P, 3-(3-Hydroxy-4-nitrobenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203994-22-3P,  
 3-[(8-Hydroxy-2,3,6,7-tetrahydro-1H,5H-benzo[ij]quinolizin-9-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203994-23-4P,  
 3-(3,5-Diisopropyl-4-hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203994-24-5P, 3-[(Benzo[b]furan-2-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203994-25-6P, 3-[[1-(4-Chlorophenyl)pyrrol-2-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203994-26-7P,  
 3-[(2-Ethylfuran-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203994-27-8P, 3-[(3,4-Dimethylthieno[2,3-b]thien-2-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203994-28-9P, 3-[(3-Bromothien-2-yl)methylidenyl]-5,7-dibromo-2-indolinone 203994-29-0P, 3-(2-Bromo-6-hydroxy-5-methoxybenzylidenyl)-5,7-dibromo-2-indolinone 203994-30-3P,  
 3-[(2-Methylfuran-5-yl)methylidenyl]-5,7-dibromo-2-indolinone 203994-31-4P, 3-[(3-Methylpyrazol-5-yl)methylidenyl]-5,7-dibromo-2-indolinone 203994-32-5P, 3-(2-Hydroxy-6-methoxy-4-methylbenzylidenyl)-5,7-dibromo-2-indolinone 203994-33-6P,  
 3-[4-(4-Formylpiperazin-1-yl)benzylidenyl]-5,7-dibromo-2-indolinone 203994-34-7P, 3-[4-(Morpholino)benzylidenyl]-5,7-dibromo-2-indolinone 203994-35-8P, 3-[[2-Chloro-4-(methoxycarbonyl)-3-(methoxycarbonyl)methyl]pyrrol-5-yl)methylidenyl]-5,7-dibromo-2-indolinone 203994-36-9P, 3-[[4-Bromo-2-(4-chlorophenyl)pyrazol-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203994-37-0P,  
 3-[(Imidazol-4-yl)methylidenyl]-5,7-dibromo-2-indolinone 203994-39-2P, 3-[(3-Bromothien-2-yl)methylidenyl]-5-iodo-2-indolinone 203994-41-6P, 3-(2-Bromo-6-hydroxy-5-methoxybenzylidenyl)-5-iodo-2-indolinone 203994-43-8P,  
 3-[(2-Methylfuran-5-yl)methylidenyl]-5-iodo-2-indolinone 203994-45-0P, 3-[(3-Methylpyrazol-5-yl)methylidenyl]-5-iodo-2-indolinone 203994-47-2P, 3-(2-Hydroxy-6-methoxy-4-methylbenzylidenyl)-5-iodo-2-indolinone 203994-49-4P,  
 3-[4-(4-Formylpiperazin-1-yl)benzylidenyl]-5-iodo-2-indolinone 203994-51-8P, 3-[4-(Morpholino)benzylidenyl]-5-iodo-2-indolinone 203994-53-0P, 3-[[2-Chloro-4-(methoxycarbonyl)-3-(methoxycarbonyl)methyl]pyrrol-5-yl)methylidenyl]-5-iodo-2-indolinone 203994-55-2P, 3-[[4-Bromo-2-(4-chlorophenyl)pyrazol-3-yl)methylidenyl]-5-iodo-2-indolinone 203994-57-4P,  
 3-[(Imidazol-4-yl)methylidenyl]-5-iodo-2-indolinone 203994-59-6P, 3-[(3-Bromothien-2-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203994-61-0P, 3-(2-Bromo-6-hydroxy-5-methoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203994-63-2P, 3-[(2-Methylfuran-5-

yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203994-65-4P,  
 3-[(3-Methylpyrazol-5-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone  
 203994-67-6P, 3-(2-Hydroxy-6-methoxy-4-methylbenzylidenyl)-5-bromo-  
 4-methyl-2-indolinone 203994-69-8P, 3-[4-(4-Formylpiperazin-1-  
 yl)benzylidenyl]-5-bromo-4-methyl-2-indolinone 203994-70-1P,  
 3-[4-(Morpholino)benzylidenyl]-5-bromo-4-methyl-2-indolinone  
 203994-72-3P, 3-[[2-Chloro-4-(methoxycarbonyl)-3-  
 [(methoxycarbonyl)methyl]pyrrol-5-yl)methylidenyl]-5-bromo-4-methyl-2-  
 indolinone 203994-74-5P, 3-[[4-Bromo-2-(4-chlorophenyl)pyrazol-3-  
 yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203994-75-6P,  
 3-[(Imidazol-4-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone  
 203994-77-8P, 3-[(3-Bromothien-2-yl)methylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203994-79-0P,  
 3-(2-Bromo-6-hydroxy-5-methoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-  
 indolinone 203994-81-4P, 3-[(2-Methylfuran-5-yl)methylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203994-83-6P,  
 3-[(3-Methylpyrazol-5-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-  
 indolinone 203994-85-8P, 3-(2-Hydroxy-6-methoxy-4-  
 methylbenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone  
 203994-87-0P, 3-[4-(4-Formylpiperazin-1-yl)benzylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203994-89-2P,  
 3-[4-(Morpholino)benzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone  
 203994-91-6P, 3-[[2-Chloro-4-(methoxycarbonyl)-3-  
 [(methoxycarbonyl)methyl]pyrrol-5-yl)methylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203994-93-8P,  
 3-[[4-Bromo-2-(4-chlorophenyl)pyrazol-3-yl)methylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203994-95-0P,  
 3-[(Imidazol-4-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone  
 203994-97-2P, 3-[(3-Bromothien-2-yl)methylidenyl]-5-[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203994-99-4P  
 , 3-(2-Bromo-6-hydroxy-5-methoxybenzylidenyl)-5-[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203995-01-1P  
 , 3-[(2-Methylfuran-5-yl)methylidenyl]-5-[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203995-03-3P  
 , 3-[(3-Methylpyrazol-5-yl)methylidenyl]-5-[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203995-05-5P  
 , 3-(2-Hydroxy-6-methoxy-4-methylbenzylidenyl)-5-[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203995-07-7P  
 , 3-[4-(4-Formylpiperazin-1-yl)benzylidenyl]-5-[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203995-09-9P  
 , 3-[4-(Morpholino)benzylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulf  
 onyl]-2-indolinone 203995-11-3P, 3-[[2-Chloro-4-  
 (methoxycarbonyl)-3-[(methoxycarbonyl)methyl]pyrrol-5-yl)methylidenyl]-5-  
 [[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone  
 203995-13-5P, 3-[[4-Bromo-2-(4-chlorophenyl)pyrazol-3-  
 yl)methylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-  
 indolinone 203995-15-7P, 3-[(Imidazol-4-yl)methylidenyl]-5-[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203995-17-9P  
 , 3-[(3-Bromothien-2-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone  
 203995-19-1P, 3-(2-Bromo-6-hydroxy-5-methoxybenzylidenyl)-5-  
 (morpholinosulfonyl)-2-indolinone 203995-21-5P,  
 3-[(2-Methylfuran-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone  
 203995-22-6P, 3-[(3-Methylpyrazol-5-yl)methylidenyl]-5-  
 (morpholinosulfonyl)-2-indolinone 203995-23-7P,  
 3-(2-Hydroxy-6-methoxy-4-methylbenzylidenyl)-5-(morpholinosulfonyl)-2-  
 indolinone 203995-24-8P, 3-[4-(4-Formylpiperazin-1-  
 yl)benzylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203995-25-9P  
 , 3-[4-(Morpholino)benzylidenyl]-5-(morpholinosulfonyl)-2-indolinone  
 203995-26-0P, 3-[[2-Chloro-4-(methoxycarbonyl)-3-

[(methoxycarbonyl)methyl]pyrrol-5-yl)methylidenyl]-5-(morpholinosulfonyl)-  
 2-indolinone 203995-27-1P, 3-[[4-Bromo-2-(4-chlorophenyl)pyrazol-  
 3-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone  
 203995-28-2P, 3-[(Imidazol-4-yl)methylidenyl]-5-  
 (morpholinosulfonyl)-2-indolinone 203995-29-3P,  
 3-[(3-Bromothien-2-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone  
 203995-30-6P, 3-(2-Bromo-6-hydroxy-5-methoxybenzylidenyl)-5-(2-  
 chloroethyl)-2-indolinone 203995-31-7P, 3-[(2-Methylfuran-5-  
 yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203995-32-8P,  
 3-[(3-Methylpyrazol-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone  
 203995-33-9P, 3-(2-Hydroxy-6-methoxy-4-methylbenzylidenyl)-5-(2-  
 chloroethyl)-2-indolinone 203995-34-0P, 3-[4-(4-Formylpiperazin-  
 1-yl)benzylidenyl]-5-(2-chloroethyl)-2-indolinone 203995-35-1P,  
 3-[4-(Morpholino)benzylidenyl]-5-(2-chloroethyl)-2-indolinone  
 203995-36-2P, 3-[[2-Chloro-4-(methoxycarbonyl)-3-  
 [(methoxycarbonyl)methyl]pyrrol-5-yl)methylidenyl]-5-(2-chloroethyl)-2-  
 indolinone 203995-37-3P, 3-[[4-Bromo-2-(4-chlorophenyl)pyrazol-3-  
 yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203995-38-4P,  
 3-[(Imidazol-4-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone  
 203995-39-5P, 3-[[2-(Ethoxycarbonyl)-4-(methoxycarbonyl)-3-  
 methylpyrrol-5-yl)methylidenyl]-5,7-dibromo-2-indolinone  
 203995-40-8P, 3-(3-tert-Butyl-4-hydroxy-5-methylbenzylidenyl)-5,7-  
 dibromo-2-indolinone 203995-41-9P, 3-[(2-Bromofuran-5-  
 yl)methylidenyl]-5,7-dibromo-2-indolinone 203995-42-0P,  
 3-[(1,3-Dimethylpyrrol-4-yl)methylidenyl]-5,7-dibromo-2-indolinone  
 203995-43-1P, 3-[(5,8-Dihydroxy-1,2,3,4-tetrahydronaphth-6-  
 yl)methylidenyl]-5,7-dibromo-2-indolinone 203995-44-2P  
 203995-45-3P 203995-46-4P, 3-[(2-Ethylthien-5-  
 yl)methylidenyl]-5,7-dibromo-2-indolinone 203995-47-5P,  
 3-(4-Methoxybenzylidenyl)-5,7-dibromo-2-indolinone 203995-48-6P,  
 3-[[2-(Ethoxycarbonyl)-4-(methoxycarbonyl)-3-methylpyrrol-5-  
 yl)methylidenyl]-5-iodo-2-indolinone 203995-49-7P,  
 3-(3-tert-Butyl-4-hydroxy-5-methylbenzylidenyl)-5-iodo-2-indolinone  
 203995-50-0P, 3-[(2-Bromofuran-5-yl)methylidenyl]-5-iodo-2-  
 indolinone 203995-51-1P, 3-[(1,3-Dimethylpyrrol-4-  
 yl)methylidenyl]-5-iodo-2-indolinone 203995-52-2P,  
 3-[(5,8-Dihydroxy-1,2,3,4-tetrahydronaphth-6-yl)methylidenyl]-5-iodo-2-  
 indolinone 203995-53-3P 203995-54-4P  
 203995-55-5P  
 , 3-[(2-Ethylthien-5-yl)methylidenyl]-5-iodo-2-indolinone 203995-56-6P  
 , 3-(4-Methoxybenzylidenyl)-5-iodo-2-indolinone 203995-57-7P,  
 3-[[2-(Ethoxycarbonyl)-4-(methoxycarbonyl)-3-methylpyrrol-5-  
 yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203995-58-8P,  
 3-(3-tert-Butyl-4-hydroxy-5-methylbenzylidenyl)-5-bromo-4-methyl-2-  
 indolinone 203995-59-9P, 3-[(2-Bromofuran-5-yl)methylidenyl]-5-  
 bromo-4-methyl-2-indolinone 203995-60-2P, 3-[(1,3-Dimethylpyrrol-  
 4-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203995-61-3P,  
 3-[(5,8-Dihydroxy-1,2,3,4-tetrahydronaphth-6-yl)methylidenyl]-5-bromo-4-  
 methyl-2-indolinone 203995-62-4P 203995-63-5P  
 203995-64-6P, 3-[(2-Ethylthien-5-yl)methylidenyl]-5-bromo-4-methyl-  
 2-indolinone 203995-65-7P, 3-(4-Methoxybenzylidenyl)-5-bromo-4-  
 methyl-2-indolinone 203995-66-8P, 3-[[2-(Ethoxycarbonyl)-4-  
 (methoxycarbonyl)-3-methylpyrrol-5-yl)methylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203995-67-9P,  
 3-(3-tert-Butyl-4-hydroxy-5-methylbenzylidenyl)-5-[(methylamino)sulfonyl]-  
 2-indolinone 203995-68-0P, 3-[(2-Bromofuran-5-yl)methylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203995-69-1P,  
 3-[(1,3-Dimethylpyrrol-4-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-  
 indolinone 203995-70-4P, 3-[(5,8-Dihydroxy-1,2,3,4-

tetrahydronaphth-6-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone  
 203995-71-5P 203995-72-6P 203995-73-7P,  
 3-[(2-Ethylthien-5-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone  
 203995-74-8P, 3-(4-Methoxybenzylidenyl)-5-[(methylamino)sulfonyl]-  
 2-indolinone 203995-75-9P, 3-[[2-(Ethoxycarbonyl)-4-  
 (methoxycarbonyl)-3-methylpyrrol-5-yl)methylidenyl]-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203995-76-0P  
 , 3-(3-tert-Butyl-4-hydroxy-5-methylbenzylidenyl)-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203995-77-1P  
 , 3-[(2-Bromofuran-5-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]  
 ]sulfonyl]-2-indolinone 203995-78-2P, 3-[(1,3-Dimethylpyrrol-4-  
 yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-  
 indolinone 203995-79-3P, 3-[(5,8-Dihydroxy-1,2,3,4-  
 tetrahydronaphth-6-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]s  
 ulfonyl]-2-indolinone 203995-80-6P 203995-81-7P  
 203995-82-8P, 3-[(2-Ethylthien-5-yl)methylidenyl]-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203995-83-9P  
 , 3-(4-Methoxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-  
 2-indolinone 203995-84-0P, 3-[[2-(Ethoxycarbonyl)-4-  
 (methoxycarbonyl)-3-methylpyrrol-5-yl)methylidenyl]-5-(morpholinosulfonyl)-  
 2-indolinone 203995-85-1P, 3-(3-tert-Butyl-4-hydroxy-5-  
 methylbenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone  
 203995-86-2P, 3-[(2-Bromofuran-5-yl)methylidenyl]-5-  
 (morpholinosulfonyl)-2-indolinone 203995-87-3P,  
 3-[(1,3-Dimethylpyrrol-4-yl)methylidenyl]-5-(morpholinosulfonyl)-2-  
 indolinone 203995-88-4P, 3-[(5,8-Dihydroxy-1,2,3,4-  
 tetrahydronaphth-6-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone  
 203995-89-5P 203995-90-8P 203995-91-9P,  
 3-[(2-Ethylthien-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone  
 203995-92-0P, 3-(4-Methoxybenzylidenyl)-5-(morpholinosulfonyl)-2-  
 indolinone 203995-93-1P, 3-[[2-(Ethoxycarbonyl)-4-  
 (methoxycarbonyl)-3-methylpyrrol-5-yl)methylidenyl]-5-(2-chloroethyl)-2-  
 indolinone 203995-94-2P, 3-(3-tert-Butyl-4-hydroxy-5-  
 methylbenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203995-95-3P,  
 3-[(2-Bromofuran-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone  
 203995-96-4P, 3-[(1,3-Dimethylpyrrol-4-yl)methylidenyl]-5-(2-  
 chloroethyl)-2-indolinone 203995-97-5P, 3-[(5,8-Dihydroxy-  
 1,2,3,4-tetrahydronaphth-6-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone  
 203995-98-6P 203995-99-7P 203996-00-3P,  
 3-[(2-Ethylthien-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone  
 203996-01-4P, 3-(4-Methoxybenzylidenyl)-5-(2-chloroethyl)-2-  
 indolinone 203996-02-5P, 3-[4-(Diethylamino)benzylidenyl]-5,7-  
 dibromo-2-indolinone 203996-03-6P, 3-[(2,4-Diethylpyrrol-5-  
 yl)methylidenyl]-5,7-dibromo-2-indolinone 203996-04-7P,  
 3-(3-Bromo-5-chloro-2-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone  
 203996-05-8P, 3-[2-[(4-Chlorophenyl)thio]benzylidenyl]-5,7-dibromo-  
 2-indolinone 203996-06-9P, 3-[(5-Chlorobenzodioxolan-6-  
 yl)methylidenyl]-5,7-dibromo-2-indolinone 203996-07-0P,  
 3-[(1,4-Benzopyranon-3-yl)methylidenyl]-5,7-dibromo-2-indolinone  
 203996-08-1P, 3-(3-Cyanobenzylidenyl)-5,7-dibromo-2-indolinone  
 203996-09-2P, 3-(4-Cyanobenzylidenyl)-5,7-dibromo-2-indolinone  
 203996-10-5P, 3-(2,5-Dihydroxybenzylidenyl)-5,7-dibromo-2-  
 indolinone 203996-11-6P, 3-(2,3-Dimethoxybenzylidenyl)-5,7-  
 dibromo-2-indolinone 203996-12-7P, 3-[4-  
 (Diethylamino)benzylidenyl]-5-iodo-2-indolinone 203996-13-8P,  
 3-[(2,4-Diethylpyrrol-5-yl)methylidenyl]-5-iodo-2-indolinone  
 203996-14-9P, 3-(3-Bromo-5-chloro-2-hydroxybenzylidenyl)-5-iodo-2-  
 indolinone 203996-15-0P, 3-[2-[(4-Chlorophenyl)thio]benzylidenyl  
 ]-5-iodo-2-indolinone

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. and testing of indolinone combinatorial library as protein kinase inhibitors)

IT 203996-16-1P, 3-[(5-Chlorobenzodioxolan-6-yl)methylidenyl]-5-iodo-2-indolinone 203996-17-2P, 3-[(1,4-Benzopyranon-3-yl)methylidenyl]-5-iodo-2-indolinone 203996-18-3P, 3-(3-Cyanobenzylidenyl)-5-iodo-2-indolinone 203996-19-4P, 3-(4-Cyanobenzylidenyl)-5-iodo-2-indolinone 203996-20-7P, 3-(2,5-Dihydroxybenzylidenyl)-5-iodo-2-indolinone 203996-21-8P, 3-(2,3-Dimethoxybenzylidenyl)-5-iodo-2-indolinone 203996-22-9P, 3-[4-(Diethylamino)benzylidenyl]-5-bromo-4-methyl-2-indolinone 203996-23-0P, 3-[(2,4-Diethylpyrrol-5-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203996-24-1P, 3-(3-Bromo-5-chloro-2-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203996-25-2P, 3-[2-[(4-Chlorophenyl)thio]benzylidenyl]-5-bromo-4-methyl-2-indolinone 203996-26-3P, 3-[(5-Chlorobenzodioxolan-6-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203996-27-4P, 3-[(1,4-Benzopyranon-3-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203996-28-5P, 3-(3-Cyanobenzylidenyl)-5-bromo-4-methyl-2-indolinone 203996-29-6P, 3-(4-Cyanobenzylidenyl)-5-bromo-4-methyl-2-indolinone 203996-30-9P, 3-(2,5-Dihydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203996-31-0P, 3-(2,3-Dimethoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203996-32-1P, 3-[4-(Diethylamino)benzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203996-33-2P, 3-[(2,4-Diethylpyrrol-5-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203996-34-3P, 3-(3-Bromo-5-chloro-2-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203996-35-4P, 3-[2-[(4-Chlorophenyl)thio]benzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203996-36-5P, 3-[(5-Chlorobenzodioxolan-6-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203996-37-6P, 3-[(1,4-Benzopyranon-3-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203996-38-7P, 3-(3-Cyanobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203996-39-8P, 3-(4-Cyanobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203996-40-1P, 3-(2,5-Dihydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203996-41-2P, 3-(2,3-Dimethoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203996-42-3P, 3-[4-(Diethylamino)benzylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203996-43-4P, 3-[(2,4-Diethylpyrrol-5-yl)methylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203996-44-5P, 3-(3-Bromo-5-chloro-2-hydroxybenzylidenyl)-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203996-45-6P, 3-[2-[(4-Chlorophenyl)thio]benzylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203996-46-7P, 3-[(5-Chlorobenzodioxolan-6-yl)methylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203996-47-8P, 3-[(1,4-Benzopyranon-3-yl)methylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203996-48-9P, 3-(3-Cyanobenzylidenyl)-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203996-49-0P, 3-(4-Cyanobenzylidenyl)-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203996-50-3P, 3-(2,5-Dihydroxybenzylidenyl)-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203996-51-4P, 3-(2,3-Dimethoxybenzylidenyl)-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203996-52-5P, 3-[4-(Diethylamino)benzylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203996-53-6P,

3-[(2,4-Diethylpyrrol-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone **203996-54-7P**, 3-(3-Bromo-5-chloro-2-hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203996-55-8P**, 3-[2-[(4-Chlorophenyl)thio]benzylidenyl]-5-(morpholinosulfonyl)-2-indolinone **203996-56-9P**, 3-[(5-Chlorobenzodioxolan-6-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone **203996-57-0P**, 3-[(1,4-Benzopyranon-3-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone **203996-58-1P**, 3-(3-Cyanobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203996-59-2P**, 3-(4-Cyanobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203996-60-5P**, 3-(2,5-Dihydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203996-61-6P**, 3-(2,3-Dimethoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **203996-62-7P**, 3-[4-(Diethylamino)benzylidenyl]-5-(2-chloroethyl)-2-indolinone **203996-63-8P**, 3-[(2,4-Diethylpyrrol-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone **203996-64-9P**, 3-(3-Bromo-5-chloro-2-hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203996-65-0P**, 3-[2-[(4-Chlorophenyl)thio]benzylidenyl]-5-(2-chloroethyl)-2-indolinone **203996-66-1P**, 3-[(5-Chlorobenzodioxolan-6-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone **203996-67-2P**, 3-[(1,4-Benzopyranon-3-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone **203996-68-3P**, 3-(3-Cyanobenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203996-69-4P**, 3-(4-Cyanobenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203996-70-7P**, 3-(2,5-Dihydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203996-71-8P**, 3-(2,3-Dimethoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone **203996-72-9P**, 3-(2,5-Dimethoxybenzylidenyl)-5,7-dibromo-2-indolinone **203996-73-0P**, 3-(2,6-Dimethoxybenzylidenyl)-5,7-dibromo-2-indolinone **203996-74-1P**, 3-(3,5-Dimethoxybenzylidenyl)-5,7-dibromo-2-indolinone **203996-75-2P**, 3-[4-(Dimethylamino)-2-methoxybenzylidenyl]-5,7-dibromo-2-indolinone **203996-76-3P**, 3-[(Fluoren-2-yl)methylidenyl]-5,7-dibromo-2-indolinone **203996-77-4P**, 3-[2-Fluoro-3-(trifluoromethyl)benzylidenyl]-5,7-dibromo-2-indolinone **203996-78-5P**, 3-[2-Fluoro-5-(trifluoromethyl)benzylidenyl]-5,7-dibromo-2-indolinone **203996-79-6P**, 3-[2-Fluoro-6-(trifluoromethyl)benzylidenyl]-5,7-dibromo-2-indolinone **203996-80-9P**, 3-[2-(Carboxymethoxy)benzylidenyl]-5,7-dibromo-2-indolinone **203996-81-0P**, 3-[(4-Methoxybenzodioxolan-6-yl)methylidenyl]-5,7-dibromo-2-indolinone **203996-82-1P**, 3-(2,5-Dimethoxybenzylidenyl)-5-iodo-2-indolinone **203996-83-2P**, 3-(2,6-Dimethoxybenzylidenyl)-5-iodo-2-indolinone **203996-84-3P**, 3-(3,5-Dimethoxybenzylidenyl)-5-iodo-2-indolinone **203996-85-4P**, 3-[4-(Dimethylamino)-2-methoxybenzylidenyl]-5-iodo-2-indolinone **203996-86-5P**, 3-[(Fluoren-2-yl)methylidenyl]-5-iodo-2-indolinone **203996-87-6P**, 3-[2-Fluoro-3-(trifluoromethyl)benzylidenyl]-5-iodo-2-indolinone **203996-88-7P**, 3-[2-Fluoro-5-(trifluoromethyl)benzylidenyl]-5-iodo-2-indolinone **203996-89-8P**, 3-[2-Fluoro-6-(trifluoromethyl)benzylidenyl]-5-iodo-2-indolinone **203996-90-1P**, 3-[2-(Carboxymethoxy)benzylidenyl]-5-iodo-2-indolinone **203996-91-2P**, 3-[(4-Methoxybenzodioxolan-6-yl)methylidenyl]-5-iodo-2-indolinone **203996-92-3P**, 3-(2,5-Dimethoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone **203996-93-4P**, 3-(2,6-Dimethoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone **203996-94-5P**, 3-(3,5-Dimethoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone **203996-95-6P**, 3-[4-(Dimethylamino)-2-methoxybenzylidenyl]-5-bromo-4-methyl-2-indolinone **203996-96-7P**, 3-[(Fluoren-2-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone **203996-97-8P**,

3-[2-Fluoro-3-(trifluoromethyl)benzylidenyl]-5-bromo-4-methyl-2-indolinone  
 203996-98-9P, 3-[2-Fluoro-5-(trifluoromethyl)benzylidenyl]-5-bromo-  
 4-methyl-2-indolinone 203996-99-0P, 3-[2-Fluoro-6-  
 (trifluoromethyl)benzylidenyl]-5-bromo-4-methyl-2-indolinone  
 203997-00-6P, 3-[2-(Carboxymethoxy)benzylidenyl]-5-bromo-4-methyl-  
 2-indolinone 203997-01-7P, 3-[(4-Methoxybenzodioxolan-6-  
 yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203997-02-8P,  
 3-(2,5-Dimethoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone  
 203997-03-9P, 3-(2,6-Dimethoxybenzylidenyl)-5-  
 [(methylamino)sulfonyl]-2-indolinone 203997-04-0P,  
 3-(3,5-Dimethoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone  
 203997-05-1P, 3-[4-(Dimethylamino)-2-methoxybenzylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203997-06-2P,  
 3-[(Fluoren-2-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone  
 203997-07-3P, 3-[2-Fluoro-3-(trifluoromethyl)benzylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203997-08-4P,  
 3-[2-Fluoro-5-(trifluoromethyl)benzylidenyl]-5-[(methylamino)sulfonyl]-2-  
 indolinone 203997-09-5P, 3-[2-Fluoro-6-  
 (trifluoromethyl)benzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone  
 203997-10-8P, 3-[2-(Carboxymethoxy)benzylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203997-11-9P,  
 3-[(4-Methoxybenzodioxolan-6-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-  
 indolinone 203997-12-0P, 3-(2,5-Dimethoxybenzylidenyl)-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203997-13-1P  
 , 3-(2,6-Dimethoxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfo  
 nyl]-2-indolinone 203997-14-2P, 3-(3,5-Dimethoxybenzylidenyl)-5-  
 [[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone  
 203997-15-3P, 3-[4-(Dimethylamino)-2-methoxybenzylidenyl]-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203997-16-4P  
 , 3-[(Fluoren-2-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulf  
 onyl]-2-indolinone 203997-17-5P, 3-[2-Fluoro-3-  
 (trifluoromethyl)benzylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone  
 203997-18-6P, 3-[2-Fluoro-5-  
 (trifluoromethyl)benzylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone  
 203997-19-7P, 3-[2-Fluoro-6-  
 (trifluoromethyl)benzylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone  
 203997-20-0P, 3-[2-  
 (Carboxymethoxy)benzylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone  
 203997-21-1P, 3-[(4-Methoxybenzodioxolan-6-  
 yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-  
 indolinone 203997-22-2P, 3-(2,5-Dimethoxybenzylidenyl)-5-  
 (morpholinosulfonyl)-2-indolinone 203997-23-3P,  
 3-(2,6-Dimethoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone  
 203997-24-4P, 3-(3,5-Dimethoxybenzylidenyl)-5-(morpholinosulfonyl)-  
 2-indolinone 203997-25-5P, 3-[4-(Dimethylamino)-2-  
 methoxybenzylidenyl]-5-(morpholinosulfonyl)-2-indolinone  
 203997-26-6P, 3-[(Fluoren-2-yl)methylidenyl]-5-  
 (morpholinosulfonyl)-2-indolinone 203997-27-7P,  
 3-[2-Fluoro-3-(trifluoromethyl)benzylidenyl]-5-(morpholinosulfonyl)-2-  
 indolinone 203997-28-8P, 3-[2-Fluoro-5-  
 (trifluoromethyl)benzylidenyl]-5-(morpholinosulfonyl)-2-indolinone  
 203997-29-9P, 3-[2-Fluoro-6-(trifluoromethyl)benzylidenyl]-5-  
 (morpholinosulfonyl)-2-indolinone 203997-30-2P,  
 3-[2-(Carboxymethoxy)benzylidenyl]-5-(morpholinosulfonyl)-2-indolinone  
 203997-31-3P, 3-[(4-Methoxybenzodioxolan-6-yl)methylidenyl]-5-  
 (morpholinosulfonyl)-2-indolinone 203997-32-4P,  
 3-(2,5-Dimethoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone  
 203997-33-5P, 3-(2,6-Dimethoxybenzylidenyl)-5-(2-chloroethyl)-2-  
 indolinone 203997-34-6P, 3-(3,5-Dimethoxybenzylidenyl)-5-(2-



chloroethyl)-2-indolinone 203997-35-7P, 3-[4-(Dimethylamino)-2-methoxybenzylidenyl]-5-(2-chloroethyl)-2-indolinone 203997-36-8P, 3-[(Fluoren-2-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203997-37-9P, 3-[2-Fluoro-3-(trifluoromethyl)benzylidenyl]-5-(2-chloroethyl)-2-indolinone 203997-38-0P, 3-[2-Fluoro-5-(trifluoromethyl)benzylidenyl]-5-(2-chloroethyl)-2-indolinone 203997-39-1P, 3-[2-Fluoro-6-(trifluoromethyl)benzylidenyl]-5-(2-chloroethyl)-2-indolinone 203997-40-4P, 3-[2-(Carboxymethoxy)benzylidenyl]-5-(2-chloroethyl)-2-indolinone 203997-41-5P, 3-[(4-Methoxybenzodioxolan-6-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203997-42-6P, 3-[(2-Methoxynaphth-1-yl)methylidenyl]-5,7-dibromo-2-indolinone 203997-43-7P, 3-[(1-Methoxynaphth-4-yl)methylidenyl]-5,7-dibromo-2-indolinone 203997-44-8P, 3-[4-(Methylthio)benzylidenyl]-5,7-dibromo-2-indolinone 203997-45-9P, 3-[(3-Methylthien-2-yl)methylidenyl]-5,7-dibromo-2-indolinone 203997-46-0P, 3-(3-Phenoxybenzylidenyl)-5,7-dibromo-2-indolinone 203997-47-1P, 3-[(Pyrid-2-yl)methylidenyl]-5,7-dibromo-2-indolinone 203997-48-2P, 3-[(Pyrid-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203997-49-3P, 3-[(Pyrid-4-yl)methylidenyl]-5,7-dibromo-2-indolinone 203997-50-6P, 3-[4-(Pyrrolidin-1-yl)benzylidenyl]-5,7-dibromo-2-indolinone 203997-51-7P, 3-[(Cyclohexen-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203997-52-8P, 3-[(2-Methoxynaphth-1-yl)methylidenyl]-5-iodo-2-indolinone 203997-53-9P, 3-[(1-Methoxynaphth-4-yl)methylidenyl]-5-iodo-2-indolinone 203997-54-0P, 3-[4-(Methylthio)benzylidenyl]-5-iodo-2-indolinone 203997-55-1P, 3-[(3-Methylthien-2-yl)methylidenyl]-5-iodo-2-indolinone 203997-56-2P, 3-(3-Phenoxybenzylidenyl)-5-iodo-2-indolinone 203997-57-3P, 3-[(Pyrid-2-yl)methylidenyl]-5-iodo-2-indolinone 203997-58-4P, 3-[(Pyrid-3-yl)methylidenyl]-5-iodo-2-indolinone 203997-59-5P, 3-[(Pyrid-4-yl)methylidenyl]-5-iodo-2-indolinone 203997-60-8P, 3-[4-(Pyrrolidin-1-yl)benzylidenyl]-5-iodo-2-indolinone 203997-61-9P, 3-[(Cyclohexen-3-yl)methylidenyl]-5-iodo-2-indolinone 203997-62-0P, 3-[(2-Methoxynaphth-1-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203997-63-1P, 3-[(1-Methoxynaphth-4-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203997-64-2P, 3-[4-(Methylthio)benzylidenyl]-5-bromo-4-methyl-2-indolinone 203997-65-3P, 3-[(3-Methylthien-2-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203997-66-4P, 3-(3-Phenoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203997-67-5P, 3-[(Pyrid-2-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203997-68-6P, 3-[(Pyrid-3-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203997-69-7P, 3-[(Pyrid-4-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203997-70-0P, 3-[4-(Pyrrolidin-1-yl)benzylidenyl]-5-bromo-4-methyl-2-indolinone 203997-71-1P, 3-[(Cyclohexen-3-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203997-72-2P, 3-[(2-Methoxynaphth-1-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203997-73-3P, 3-[(1-Methoxynaphth-4-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203997-74-4P, 3-[4-(Methylthio)benzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203997-75-5P, 3-[(3-Methylthien-2-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203997-76-6P, 3-(3-Phenoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 203997-77-7P, 3-[(Pyrid-2-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203997-78-8P, 3-[(Pyrid-3-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203997-79-9P, 3-[(Pyrid-4-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 203997-81-3P, 3-[4-(Pyrrolidin-1-yl)benzylidenyl]-5-

[(methylamino)sulfonyl]-2-indolinone 203997-83-5P,  
 3-[(Cyclohexen-3-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone  
 203997-85-7P, 3-[(2-Methoxynaphth-1-yl)methylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203997-87-9P  
 , 3-[(1-Methoxynaphth-4-yl)methylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203997-89-1P  
 , 3-[4-(Methylthio)benzylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203997-90-4P, 3-[(3-Methylthien-2-yl)methylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203997-91-5P, 3-(3-Phenoxybenzylidenyl)-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203997-92-6P  
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 , 3-[4-(Pyrrolidin-1-yl)benzylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203997-96-0P, 3-[(Cyclohexen-3-yl)methylidenyl]-5-[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203997-97-1P, 3-[(2-Methoxynaphth-1-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203997-98-2P,  
 3-[(1-Methoxynaphth-4-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203997-99-3P, 3-[4-(Methylthio)benzylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203998-00-9P,  
 3-[(3-Methylthien-2-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203998-01-0P, 3-(3-Phenoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203998-02-1P, 3-[(Pyrid-2-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203998-03-2P,  
 3-[(Pyrid-3-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203998-04-3P, 3-[(Pyrid-4-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203998-05-4P, 3-[4-(Pyrrolidin-1-yl)benzylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203998-06-5P,  
 3-[(Cyclohexen-3-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203998-07-6P, 3-[(2-Methoxynaphth-1-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-08-7P, 3-[(1-Methoxynaphth-4-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-09-8P,  
 3-[4-(Methylthio)benzylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-10-1P, 3-[(3-Methylthien-2-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-11-2P, 3-(3-Phenoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203998-12-3P  
 , 3-[(Pyrid-2-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-13-4P, 3-[(Pyrid-3-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-14-5P, 3-[(Pyrid-4-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-15-6P, 3-[4-(Pyrrolidin-1-yl)benzylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-16-7P,  
 3-[(Cyclohexen-3-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-17-8P, 3-(2,3,4-Trimethoxybenzylidenyl)-5,7-dibromo-2-indolinone 203998-18-9P, 3-(2,4,5-Trimethoxybenzylidenyl)-5,7-dibromo-2-indolinone 203998-19-0P, 3-(3,4,5-Trimethoxybenzylidenyl)-5,7-dibromo-2-indolinone 203998-20-3P,  
 3-[(1-Acetylindol-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-21-4P, 3-[(6-Chloro-1,4-benzopyranon-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-22-5P  
 , 3-[[2-(2-Chlorophenyl)furan-5-yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-23-6P, 3-[[2-Chloroquinolin-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-24-7P, 3-[(6,8-Dibromo-1,4-benzopyranon-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-25-8P,  
 3-[(2,5-Dimethoxytetrahydrofuran-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-26-9P, 3-[(2,3-Dimethylfuran-5-

yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-27-0P,  
 3-(2,3,4-Trimethoxybenzylidenyl)-5-iodo-2-indolinone 203998-28-1P  
 , 3-(2,4,5-Trimethoxybenzylidenyl)-5-iodo-2-indolinone  
 203998-29-2P, 3-(3,4,5-Trimethoxybenzylidenyl)-5-iodo-2-indolinone  
 203998-30-5P, 3-[(1-Acetylindol-3-yl)methylidenyl]-5-iodo-2-  
 indolinone 203998-31-6P, 3-[(6-Chloro-1,4-benzopyranon-3-  
 yl)methylidenyl]-5-iodo-2-indolinone 203998-32-7P,  
 3-[[2-(2-Chlorophenyl)furan-5-yl]methylidenyl]-5-iodo-2-indolinone  
 203998-33-8P, 3-[(2-Chloroquinolin-3-yl)methylidenyl]-5-iodo-2-  
 indolinone 203998-34-9P, 3-[(6,8-Dibromo-1,4-benzopyranon-3-  
 yl)methylidenyl]-5-iodo-2-indolinone 203998-35-0P,  
 3-[(2,5-Dimethoxytetrahydrofuran-3-yl)methylidenyl]-5-iodo-2-indolinone  
 203998-36-1P, 3-[(2,3-Dimethylfuran-5-yl)methylidenyl]-5-iodo-2-  
 indolinone 203998-37-2P, 3-(2,3,4-Trimethoxybenzylidenyl)-5-  
 bromo-4-methyl-2-indolinone 203998-38-3P, 3-(2,4,5-  
 Trimethoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 203998-39-4P  
 , 3-(3,4,5-Trimethoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone  
 203998-40-7P, 3-[(1-Acetylindol-3-yl)methylidenyl]-5-bromo-4-  
 methyl-2-indolinone 203998-41-8P, 3-[(6-Chloro-1,4-benzopyranon-  
 3-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203998-42-9P,  
 3-[[2-(2-Chlorophenyl)furan-5-yl]methylidenyl]-5-bromo-4-methyl-2-  
 indolinone 203998-43-0P, 3-[(2-Chloroquinolin-3-yl)methylidenyl]-  
 5-bromo-4-methyl-2-indolinone 203998-44-1P, 3-[(6,8-Dibromo-1,4-  
 benzopyranon-3-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone  
 203998-45-2P, 3-[(2,5-Dimethoxytetrahydrofuran-3-yl)methylidenyl]-  
 5-bromo-4-methyl-2-indolinone 203998-46-3P, 3-[(2,3-  
 Dimethylfuran-5-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone  
 203998-47-4P, 3-(2,3,4-Trimethoxybenzylidenyl)-5-  
 [(methylamino)sulfonyl]-2-indolinone 203998-48-5P,  
 3-(2,4,5-Trimethoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone  
 203998-49-6P, 3-(3,4,5-Trimethoxybenzylidenyl)-5-  
 [(methylamino)sulfonyl]-2-indolinone 203998-50-9P,  
 3-[(1-Acetylindol-3-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-  
 indolinone 203998-51-0P, 3-[(6-Chloro-1,4-benzopyranon-3-  
 yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone  
 203998-52-1P, 3-[[2-(2-Chlorophenyl)furan-5-yl]methylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203998-53-2P,  
 3-[(2-Chloroquinolin-3-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-  
 indolinone 203998-54-3P, 3-[(6,8-Dibromo-1,4-benzopyranon-3-  
 yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone  
 203998-55-4P, 3-[(2,5-Dimethoxytetrahydrofuran-3-yl)methylidenyl]-  
 5-[(methylamino)sulfonyl]-2-indolinone  
 RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic  
 preparation); THU (Therapeutic use); BIOL (Biological study); PREP  
 (Preparation); USES (Uses)

(prepn. and testing of indolinone combinatorial library as protein  
 kinase inhibitors)

IT 203998-56-5P, 3-[(2,3-Dimethylfuran-5-yl)methylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203998-57-6P,  
 3-(2,3,4-Trimethoxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulf  
 onyl]-2-indolinone 203998-58-7P, 3-(2,4,5-  
 Trimethoxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-  
 indolinone 203998-59-8P, 3-(3,4,5-Trimethoxybenzylidenyl)-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203998-60-1P  
 , 3-[(1-Acetylindol-3-yl)methylidenyl]-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203998-61-2P  
 , 3-[(6-Chloro-1,4-benzopyranon-3-yl)methylidenyl]-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203998-62-3P  
 , 3-[[2-(2-Chlorophenyl)furan-5-yl]methylidenyl]-5-[[[4-

(trifluoromethyl)phenyl]amino)sulfonyl]-2-indolinone 203998-63-4P  
 , 3-[(2-Chloroquinolin-3-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino)sulfonyl]-2-indolinone 203998-64-5P  
 , 3-[(6,8-Dibromo-1,4-benzopyranon-3-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino)sulfonyl]-2-indolinone 203998-65-6P  
 , 3-[(2,5-Dimethoxytetrahydrofuran-3-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino)sulfonyl]-2-indolinone 203998-66-7P  
 , 3-[(2,3-Dimethylfuran-5-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino)sulfonyl]-2-indolinone 203998-67-8P\*\*\*,  
 3-(2,3,4-Trimethoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone  
 \*\*\*203998-68-9P, 3-(2,4,5-Trimethoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203998-69-0P,  
 3-(3,4,5-Trimethoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 203998-70-3P, 3-[(1-Acetylinol-3-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203998-71-4P,  
 3-[(6-Chloro-1,4-benzopyranon-3-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203998-72-5P, 3-[[2-(2-Chlorophenyl)furan-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203998-73-6P  
 , 3-[(2-Chloroquinolin-3-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203998-74-7P, 3-[(6,8-Dibromo-1,4-benzopyranon-3-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203998-75-8P  
 , 3-[(2,5-Dimethoxytetrahydrofuran-3-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203998-76-9P,  
 3-[(2,3-Dimethylfuran-5-yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203998-77-0P, 3-(2,3,4-Trimethoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203998-78-1P, 3-(2,4,5-Trimethoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203998-79-2P, 3-(3,4,5-Trimethoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 203998-80-5P, 3-[(1-Acetylinol-3-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-81-6P, 3-[(6-Chloro-1,4-benzopyranon-3-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-82-7P, 3-[[2-(2-Chlorophenyl)furan-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-83-8P, 3-[(2-Chloroquinolin-3-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-84-9P,  
 3-[(6,8-Dibromo-1,4-benzopyranon-3-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-85-0P, 3-[(2,5-Dimethoxytetrahydrofuran-3-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-86-1P,  
 3-[(2,3-Dimethylfuran-5-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 203998-87-2P, 3-[(9-Ethylcarbazol-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-88-3P, 3-[(6,7-Dimethyl-1,4-benzopyron-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-89-4P,  
 3-[[4-(Propen-2-yl)cyclohexen-1-yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-90-7P, 3-[(6-Isopropyl-1,4-benzopyron-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-91-8P, 3-[(6-Methyl-1,4-benzopyron-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-92-9P, 3-[(6-Nitro-1,4-benzopyron-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-93-0P, 3-[(Pyrimid-2,4-dion-5-yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-94-1P,  
 3-[(5-Methoxyindol-3-yl)methylidenyl]-5,7-dibromo-2-indolinone 203998-95-2P 203998-96-3P, 3-[[2-(2-Nitrophenyl)furan-5-yl)methylidene]-5,7-dibromo-2-indolinone 203998-97-4P,  
 3-[(9-Ethylcarbazol-3-yl)methylidenyl]-5-iodo-2-indolinone 203999-01-3P, 3-[(6,7-Dimethyl-1,4-benzopyron-3-yl)methylidenyl]-5-iodo-2-indolinone 203999-02-4P, 3-[[4-(Propen-2-yl)cyclohexen-1-yl)methylidenyl]-5-iodo-2-indolinone 203999-03-5P,  
 3-[(6-Isopropyl-1,4-benzopyron-3-yl)methylidenyl]-5-iodo-2-indolinone 203999-04-6P, 3-[(6-Methyl-1,4-benzopyron-3-yl)methylidenyl]-5-iodo-2-indolinone 203999-05-7P, 3-[(6-Nitro-1,4-benzopyron-3-yl)methylidenyl]-5-iodo-2-indolinone 203999-06-8P,

3-[(Pyrimid-2,4-dion-5-yl)methylidenyl]-5-iodo-2-indolinone  
 203999-07-9P, 3-[(5-Methoxyindol-3-yl)methylidenyl]-5-iodo-2-  
 indolinone 203999-08-0P 203999-09-1P,  
 3-[[2-(2-Nitrophenyl)furan-5-yl)methylidene]-5-iodo-2-indolinone  
 203999-10-4P, 3-[(9-Ethylcarbazol-3-yl)methylidenyl]-5-bromo-4-  
 methyl-2-indolinone 203999-11-5P, 3-[(6,7-Dimethyl-1,4-  
 benzopyron-3-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone  
 203999-12-6P, 3-[[4-(Propen-2-yl)cyclohexen-1-yl)methylidenyl]-5-  
 bromo-4-methyl-2-indolinone 203999-13-7P, 3-[(6-Isopropyl-1,4-  
 benzopyron-3-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone  
 203999-14-8P, 3-[(6-Methyl-1,4-benzopyron-3-yl)methylidenyl]-5-  
 bromo-4-methyl-2-indolinone 203999-15-9P, 3-[(6-Nitro-1,4-  
 benzopyron-3-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone  
 203999-16-0P 203999-17-1P, 3-[(5-Methoxyindol-3-  
 yl)methylidenyl]-5-bromo-4-methyl-2-indolinone 203999-18-2P  
 203999-19-3P, 3-[[2-(2-Nitrophenyl)furan-5-yl)methylidene]-5-bromo-  
 4-methyl-2-indolinone 203999-20-6P, 3-[(9-Ethylcarbazol-3-  
 yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone  
 203999-21-7P, 3-[(6,7-Dimethyl-1,4-benzopyron-3-yl)methylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203999-22-8P,  
 3-[[4-(Propen-2-yl)cyclohexen-1-yl)methylidenyl]-5-[(methylamino)sulfonyl]-  
 2-indolinone 203999-23-9P, 3-[(6-Isopropyl-1,4-benzopyron-3-  
 yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone  
 203999-24-0P, 3-[(6-Methyl-1,4-benzopyron-3-yl)methylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 203999-25-1P,  
 3-[(6-Nitro-1,4-benzopyron-3-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-  
 indolinone 203999-26-2P 203999-27-3P,  
 3-[(5-Methoxyindol-3-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-  
 indolinone 203999-28-4P 203999-29-5P,  
 3-[[2-(2-Nitrophenyl)furan-5-yl)methylidene]-5-[(methylamino)sulfonyl]-2-  
 indolinone 203999-30-8P, 3-[(9-Ethylcarbazol-3-yl)methylidenyl]-  
 5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone  
 203999-31-9P, 3-[(6,7-Dimethyl-1,4-benzopyron-3-yl)methylidenyl]-5-  
 [[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone  
 203999-32-0P, 3-[[4-(Propen-2-yl)cyclohexen-1-yl)methylidenyl]-5-  
 [[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone  
 203999-33-1P, 3-[(6-Isopropyl-1,4-benzopyron-3-yl)methylidenyl]-5-  
 [[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone  
 203999-36-4P, 3-[(6-Methyl-1,4-benzopyron-3-yl)methylidenyl]-5-  
 [[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone  
 203999-38-6P, 3-[(6-Nitro-1,4-benzopyron-3-yl)methylidenyl]-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203999-40-0P  
 203999-42-2P, 3-[(5-Methoxyindol-3-yl)methylidenyl]-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203999-44-4P  
 203999-46-6P, 3-[[2-(2-Nitrophenyl)furan-5-yl)methylidene]-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 203999-47-7P  
 , 3-[(9-Ethylcarbazol-3-yl)methylidenyl]-5-(morpholinosulfonyl)-2-  
 indolinone 203999-48-8P, 3-[(6,7-Dimethyl-1,4-benzopyron-3-  
 yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203999-49-9P  
 , 3-[[4-(Propen-2-yl)cyclohexen-1-yl)methylidenyl]-5-(morpholinosulfonyl)-  
 2-indolinone 203999-50-2P, 3-[(6-Isopropyl-1,4-benzopyron-3-  
 yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203999-51-3P  
 , 3-[(6-Methyl-1,4-benzopyron-3-yl)methylidenyl]-5-(morpholinosulfonyl)-2-  
 indolinone 203999-52-4P, 3-[(6-Nitro-1,4-benzopyron-3-  
 yl)methylidenyl]-5-(morpholinosulfonyl)-2-indolinone 203999-53-5P  
 203999-54-6P, 3-[(5-Methoxyindol-3-yl)methylidenyl]-5-  
 (morpholinosulfonyl)-2-indolinone 203999-55-7P  
 203999-56-8P, 3-[[2-(2-Nitrophenyl)furan-5-yl)methylidene]-5-  
 (morpholinosulfonyl)-2-indolinone 203999-57-9P,

3-[(9-Ethylcarbazol-3-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone  
 203999-58-0P, 3-[(6,7-Dimethyl-1,4-benzopyron-3-yl)methylidenyl]-5-  
 (2-chloroethyl)-2-indolinone 203999-59-1P, 3-[4-(Propen-2-  
 yl)cyclohexen-1-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone  
 203999-61-5P, 3-[(6-Isopropyl-1,4-benzopyron-3-yl)methylidenyl]-5-  
 (2-chloroethyl)-2-indolinone 203999-63-7P, 3-[(6-Methyl-1,4-  
 benzopyron-3-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone  
 203999-65-9P, 3-[(6-Nitro-1,4-benzopyron-3-yl)methylidenyl]-5-(2-  
 chloroethyl)-2-indolinone 203999-67-1P 203999-69-3P,  
 3-[(5-Methoxyindol-3-yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone  
 203999-71-7P 203999-73-9P, 3-[[2-(2-Nitrophenyl)furan-5-  
 yl)methylidene]-5-(2-chloroethyl)-2-indolinone 203999-77-3P,  
 3-(3,5-Diisopropyl-4-methoxybenzylidenyl)-5,7-dibromo-2-indolinone  
 203999-79-5P, 3-(3,5-Diisopropyl-4-phenoxybenzylidenyl)-5,7-  
 dibromo-2-indolinone 203999-81-9P, 3-(3-tert-Butyl-4-  
 methoxybenzylidenyl)-5,7-dibromo-2-indolinone 203999-83-1P,  
 3-[4-(Benzyloxy)-3-tert-butylbenzylidenyl]-5,7-dibromo-2-indolinone  
 203999-85-3P, 3-(3-Bromo-5-tert-butyl-4-methoxybenzylidenyl)-5,7-  
 dibromo-2-indolinone 203999-87-5P, 3-[4-(Benzyloxy)-3-bromo-5-  
 tert-butylbenzylidenyl]-5,7-dibromo-2-indolinone 203999-89-7P,  
 3-(3-tert-Butyl-5-chloro-4-methoxybenzylidenyl)-5,7-dibromo-2-indolinone  
 203999-91-1P, 3-[4-(Benzyloxy)-5-tert-butyl-3-chlorobenzylidenyl]-  
 5,7-dibromo-2-indolinone 203999-93-3P, 3-(3-tert-Butyl-5-iodo-4-  
 methoxybenzylidenyl)-5,7-dibromo-2-indolinone 203999-97-7P,  
 3-(3,5-Diisopropyl-4-methoxybenzylidenyl)-5-iodo-2-indolinone  
 203999-99-9P, 3-(3,5-Diisopropyl-4-phenoxybenzylidenyl)-5-iodo-2-  
 indolinone 204000-01-1P, 3-(3-tert-Butyl-4-methoxybenzylidenyl)-  
 5-iodo-2-indolinone 204000-03-3P, 3-[4-(Benzyloxy)-3-tert-  
 butylbenzylidenyl]-5-iodo-2-indolinone 204000-05-5P,  
 3-(3-Bromo-5-tert-butyl-4-methoxybenzylidenyl)-5-iodo-2-indolinone  
 204000-07-7P, 3-[4-(Benzyloxy)-3-bromo-5-tert-butylbenzylidenyl]-5-  
 iodo-2-indolinone 204000-09-9P, 3-(3-tert-Butyl-5-chloro-4-  
 methoxybenzylidenyl)-5-iodo-2-indolinone 204000-13-5P,  
 3-[4-(Benzyloxy)-5-tert-butyl-3-chlorobenzylidenyl]-5-iodo-2-indolinone  
 204000-17-9P, 3-(3-tert-Butyl-5-iodo-4-methoxybenzylidenyl)-5-iodo-  
 2-indolinone 204000-21-5P, 3-(3,5-Diisopropyl-4-  
 methoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 204000-23-7P,  
 3-(3,5-Diisopropyl-4-phenoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone  
 204000-25-9P, 3-(3-tert-Butyl-4-methoxybenzylidenyl)-5-bromo-4-  
 methyl-2-indolinone 204000-27-1P, 3-[4-(Benzyloxy)-3-tert-  
 butylbenzylidenyl]-5-bromo-4-methyl-2-indolinone 204000-29-3P,  
 3-(3-Bromo-5-tert-butyl-4-methoxybenzylidenyl)-5-bromo-4-methyl-2-  
 indolinone 204000-31-7P, 3-[4-(Benzyloxy)-3-bromo-5-tert-  
 butylbenzylidenyl]-5-bromo-4-methyl-2-indolinone 204000-33-9P,  
 3-(3-tert-Butyl-5-chloro-4-methoxybenzylidenyl)-5-bromo-4-methyl-2-  
 indolinone 204000-34-0P, 3-[4-(Benzyloxy)-5-tert-butyl-3-  
 chlorobenzylidenyl]-5-bromo-4-methyl-2-indolinone 204000-36-2P,  
 3-(3-tert-Butyl-5-iodo-4-methoxybenzylidenyl)-5-bromo-4-methyl-2-  
 indolinone 204000-38-4P, 3-(3,5-Diisopropyl-4-  
 methoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone  
 204000-39-5P, 3-(3,5-Diisopropyl-4-phenoxybenzylidenyl)-5-  
 [(methylamino)sulfonyl]-2-indolinone 204000-40-8P,  
 3-(3-tert-Butyl-4-methoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-  
 indolinone 204000-41-9P, 3-[4-(Benzyloxy)-3-tert-  
 butylbenzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone  
 204000-42-0P, 3-(3-Bromo-5-tert-butyl-4-methoxybenzylidenyl)-5-  
 [(methylamino)sulfonyl]-2-indolinone 204000-43-1P,  
 3-[4-(Benzyloxy)-3-bromo-5-tert-butylbenzylidenyl]-5-  
 [(methylamino)sulfonyl]-2-indolinone 204000-44-2P,

3-(3-tert-Butyl-5-chloro-4-methoxybenzylidenyl)-5-[(methylamino)sulfonyl]-  
 2-indolinone **204000-45-3P**, 3-[4-(Benzyloxy)-5-tert-butyl-3-  
 chlorobenzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone  
**204000-46-4P**, 3-(3-tert-Butyl-5-iodo-4-methoxybenzylidenyl)-5-  
 [(methylamino)sulfonyl]-2-indolinone **204000-48-6P**,  
 3-(3,5-Diisopropyl-4-methoxybenzylidenyl)-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204000-49-7P**  
 , 3-(3,5-Diisopropyl-4-phenoxybenzylidenyl)-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204000-50-0P**  
 , 3-(3-tert-Butyl-4-methoxybenzylidenyl)-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204000-51-1P**  
 , 3-[4-(Benzyloxy)-3-tert-butylbenzylidenyl]-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204000-52-2P**  
 , 3-(3-Bromo-5-tert-butyl-4-methoxybenzylidenyl)-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204000-53-3P**  
 , 3-[4-(Benzyloxy)-3-bromo-5-tert-butylbenzylidenyl]-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204000-54-4P**  
 , 3-(3-tert-Butyl-5-chloro-4-methoxybenzylidenyl)-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204000-55-5P**  
 , 3-[4-(Benzyloxy)-5-tert-butyl-3-chlorobenzylidenyl]-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204000-56-6P**  
 , 3-(3-tert-Butyl-5-iodo-4-methoxybenzylidenyl)-5-[[[4-  
 (trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204000-58-8P**  
 , 3-(3,5-Diisopropyl-4-methoxybenzylidenyl)-5-(morpholinosulfonyl)-2-  
 indolinone **204000-59-9P**, 3-(3,5-Diisopropyl-4-  
 phenoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone  
**204000-60-2P**, 3-(3-tert-Butyl-4-methoxybenzylidenyl)-5-  
 (morpholinosulfonyl)-2-indolinone **204000-61-3P**,  
 3-[4-(Benzyloxy)-3-tert-butylbenzylidenyl]-5-(morpholinosulfonyl)-2-  
 indolinone **204000-62-4P**, 3-(3-Bromo-5-tert-butyl-4-  
 methoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone  
**204000-63-5P**, 3-[4-(Benzyloxy)-3-bromo-5-tert-butylbenzylidenyl]-5-  
 (morpholinosulfonyl)-2-indolinone **204000-64-6P**,  
 3-(3-tert-Butyl-5-chloro-4-methoxybenzylidenyl)-5-(morpholinosulfonyl)-2-  
 indolinone **204000-65-7P**, 3-[4-(Benzyloxy)-5-tert-butyl-3-  
 chlorobenzylidenyl]-5-(morpholinosulfonyl)-2-indolinone  
**204000-66-8P**, 3-(3-tert-Butyl-5-iodo-4-methoxybenzylidenyl)-5-  
 (morpholinosulfonyl)-2-indolinone **204000-68-0P**,  
 3-(3,5-Diisopropyl-4-methoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone  
**204000-69-1P**, 3-(3,5-Diisopropyl-4-phenoxybenzylidenyl)-5-(2-  
 chloroethyl)-2-indolinone **204000-70-4P**, 3-(3-tert-Butyl-4-  
 methoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone **204000-71-5P**  
 , 3-[4-(Benzyloxy)-3-tert-butylbenzylidenyl]-5-(2-chloroethyl)-2-  
 indolinone **204000-72-6P**, 3-(3-Bromo-5-tert-butyl-4-  
 methoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone **204000-73-7P**  
 , 3-[4-(Benzyloxy)-3-bromo-5-tert-butylbenzylidenyl]-5-(2-chloroethyl)-2-  
 indolinone **204000-74-8P**, 3-(3-tert-Butyl-5-chloro-4-  
 methoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone **204000-75-9P**  
 , 3-[4-(Benzyloxy)-5-tert-butyl-3-chlorobenzylidenyl]-5-(2-chloroethyl)-2-  
 indolinone **204000-76-0P**, 3-(3-tert-Butyl-5-iodo-4-  
 methoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone **204000-77-1P**  
 , 3-[4-(Benzyloxy)-3-tert-butyl-5-iodobenzylidenyl]-5,7-dibromo-2-  
 indolinone **204000-78-2P**, 3-(3-tert-Butyl-4-methoxy-5-  
 nitrobenzylidenyl)-5,7-dibromo-2-indolinone **204000-79-3P**,  
 3-(3,5-Di-tert-butyl-4-methoxybenzylidenyl)-5,7-dibromo-2-indolinone  
**204000-80-6P**, 3-[4-(Benzyloxy)-3,5-di-tert-butylbenzylidenyl]-5,7-  
 dibromo-2-indolinone **204000-81-7P**, 3-(3,5-Dimethyl-4-  
 methoxybenzylidenyl)-5,7-dibromo-2-indolinone **204000-82-8P**,  
 3-[4-(Benzyloxy)-3,5-dimethylbenzylidenyl]-5,7-dibromo-2-indolinone

204000-83-9P, 3-(5-Bromo-2-hydroxy-3-methoxybenzylidenyl)-5,7-dibromo-2-indolinone 204000-84-0P, 3-(5-Bromo-2-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 204000-85-1P, 3-(2-Hydroxy-5-nitrobenzylidenyl)-5,7-dibromo-2-indolinone 204000-86-2P, 3-(4-Hydroxy-3-methoxy-2-nitrobenzylidenyl)-5,7-dibromo-2-indolinone 204000-87-3P, 3-[4-(Benzyloxy)-3-tert-butyl-5-iodobenzylidenyl]-5-iodo-2-indolinone 204000-88-4P, 3-(3-tert-Butyl-4-methoxy-5-nitrobenzylidenyl)-5-iodo-2-indolinone 204000-89-5P, 3-(3,5-Di-tert-butyl-4-methoxybenzylidenyl)-5-iodo-2-indolinone 204000-90-8P, 3-[4-(Benzyloxy)-3,5-di-tert-butylbenzylidenyl]-5-iodo-2-indolinone 204000-91-9P, 3-(3,5-Dimethyl-4-methoxybenzylidenyl)-5-iodo-2-indolinone 204000-92-0P, 3-[4-(Benzyloxy)-3,5-dimethylbenzylidenyl]-5-iodo-2-indolinone 204000-93-1P, 3-(5-Bromo-2-hydroxy-3-methoxybenzylidenyl)-5-iodo-2-indolinone 204000-94-2P, 3-(5-Bromo-2-hydroxybenzylidenyl)-5-iodo-2-indolinone 204000-95-3P, 3-(2-Hydroxy-5-nitrobenzylidenyl)-5-iodo-2-indolinone 204000-96-4P, 3-(4-Hydroxy-3-methoxy-2-nitrobenzylidenyl)-5-iodo-2-indolinone 204000-97-5P, 3-[4-(Benzyloxy)-3-tert-butyl-5-iodobenzylidenyl]-5-bromo-4-methyl-2-indolinone 204000-98-6P, 3-(3-tert-Butyl-4-methoxy-5-nitrobenzylidenyl)-5-bromo-4-methyl-2-indolinone 204000-99-7P, 3-(3,5-Di-tert-butyl-4-methoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 204001-00-3P, 3-[4-(Benzyloxy)-3,5-di-tert-butylbenzylidenyl]-5-bromo-4-methyl-2-indolinone 204001-01-4P, 3-(3,5-Dimethyl-4-methoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 204001-02-5P, 3-[4-(Benzyloxy)-3,5-dimethylbenzylidenyl]-5-bromo-4-methyl-2-indolinone 204001-03-6P, 3-(5-Bromo-2-hydroxy-3-methoxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 204001-04-7P, 3-(5-Bromo-2-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone 204001-05-8P, 3-(2-Hydroxy-5-nitrobenzylidenyl)-5-bromo-4-methyl-2-indolinone 204001-06-9P, 3-(4-Hydroxy-3-methoxy-2-nitrobenzylidenyl)-5-bromo-4-methyl-2-indolinone 204001-07-0P, 3-[4-(Benzyloxy)-3-tert-butyl-5-iodobenzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 204001-08-1P, 3-(3-tert-Butyl-4-methoxy-5-nitrobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 204001-09-2P, 3-(3,5-Di-tert-butyl-4-methoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 204001-10-5P, 3-[4-(Benzyloxy)-3,5-di-tert-butylbenzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 204001-11-6P, 3-(3,5-Dimethyl-4-methoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 204001-12-7P, 3-[4-(Benzyloxy)-3,5-dimethylbenzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone 204001-13-8P, 3-(5-Bromo-2-hydroxy-3-methoxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 204001-14-9P, 3-(5-Bromo-2-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 204001-15-0P, 3-(2-Hydroxy-5-nitrobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 204001-16-1P, 3-(4-Hydroxy-3-methoxy-2-nitrobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone 204001-17-2P, 3-[4-(Benzyloxy)-3-tert-butyl-5-iodobenzylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 204001-18-3P, 3-(3-tert-Butyl-4-methoxy-5-nitrobenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 204001-19-4P, 3-(3,5-Di-tert-butyl-4-methoxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 204001-21-8P, 3-[4-(Benzyloxy)-3,5-di-tert-butylbenzylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 204001-22-9P, 3-(3,5-Dimethyl-4-methoxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone 204001-23-0P, 3-[4-(Benzyloxy)-3,5-dimethylbenzylidenyl]-5-[[[4-



(trifluoromethyl)phenyl]amino)sulfonyl]-2-indolinone 204001-24-1P,  
 3-(5-Bromo-2-hydroxy-3-methoxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino)sulfonyl]-2-indolinone 204001-25-2P,  
 3-(5-Bromo-2-hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino)sulfonyl]-2-indolinone 204001-26-3P, 3-(2-Hydroxy-5-nitrobenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino)sulfonyl]-2-indolinone 204001-27-4P,  
 3-(4-Hydroxy-3-methoxy-2-nitrobenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino)sulfonyl]-2-indolinone 204001-28-5P,  
 3-[4-(Benzyloxy)-3-tert-butyl-5-iodobenzylidenyl]-5-(morpholinosulfonyl)-2-indolinone 204001-29-6P, 3-(3-tert-Butyl-4-methoxy-5-nitrobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 204001-30-9P, 3-(3,5-Di-tert-butyl-4-methoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 204001-31-0P,  
 3-[4-(Benzyloxy)-3,5-di-tert-butylbenzylidenyl]-5-(morpholinosulfonyl)-2-indolinone 204001-32-1P, 3-(3,5-Dimethyl-4-methoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 204001-33-2P,  
 3-[4-(Benzyloxy)-3,5-dimethylbenzylidenyl]-5-(morpholinosulfonyl)-2-indolinone 204001-34-3P, 3-(5-Bromo-2-hydroxy-3-methoxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 204001-35-4P, 3-(5-Bromo-2-hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 204001-36-5P,  
 3-(2-Hydroxy-5-nitrobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 204001-37-6P, 3-(4-Hydroxy-3-methoxy-2-nitrobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone 204001-38-7P,  
 3-[4-(Benzyloxy)-3-tert-butyl-5-iodobenzylidenyl]-5-(2-chloroethyl)-2-indolinone 204001-39-8P, 3-(3-tert-Butyl-4-methoxy-5-nitrobenzylidenyl)-5-(2-chloroethyl)-2-indolinone 204001-40-1P,  
 3-(3,5-Di-tert-butyl-4-methoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 204001-41-2P, 3-[4-(Benzyloxy)-3,5-di-tert-butylbenzylidenyl]-5-(2-chloroethyl)-2-indolinone 204001-42-3P, 3-(3,5-Dimethyl-4-methoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 204001-43-4P,  
 3-[4-(Benzyloxy)-3,5-dimethylbenzylidenyl]-5-(2-chloroethyl)-2-indolinone 204001-44-5P, 3-(5-Bromo-2-hydroxy-3-methoxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 204001-45-6P,  
 3-(5-Bromo-2-hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 204001-46-7P, 3-(2-Hydroxy-5-nitrobenzylidenyl)-5-(2-chloroethyl)-2-indolinone 204001-47-8P, 3-(4-Hydroxy-3-methoxy-2-nitrobenzylidenyl)-5-(2-chloroethyl)-2-indolinone 204001-48-9P,  
 3-(3-Ethoxy-2-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone  
 RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (prepn. and testing of indolinone combinatorial library as protein kinase inhibitors)  
 IT 204001-49-0P, 3-(3,5-Dichloro-2-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 204001-50-3P, 3-(5-Chloro-2-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 204001-51-4P, 3-[4-(Diethylamino)-2-hydroxybenzylidenyl]-5,7-dibromo-2-indolinone 204001-52-5P,  
 3-(4-Nitrobenzylidenyl)-5,7-dibromo-2-indolinone 204001-53-6P, 3-(3,5-Dibromo-2-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 204001-54-7P, 3-(3-Fluoro-2-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 204001-55-8P, 3-(3-Bromo-4-hydroxybenzylidenyl)-5,7-dibromo-2-indolinone 204001-56-9P, 3-(4-tert-Butylbenzylidenyl)-5,7-dibromo-2-indolinone 204001-57-0P, 3-[(2-Bromothien-5-yl)methylidenyl]-5,7-dibromo-2-indolinone 204001-58-1P,  
 3-(3-Ethoxy-2-hydroxybenzylidenyl)-5-iodo-2-indolinone 204001-59-2P, 3-(3,5-Dichloro-2-hydroxybenzylidenyl)-5-iodo-2-indolinone 204001-60-5P, 3-(5-Chloro-2-hydroxybenzylidenyl)-5-

iodo-2-indolinone **204001-61-6P**, 3-[4-(Diethylamino)-2-hydroxybenzylidenyl]-5-iodo-2-indolinone **204001-62-7P**,  
 3-(4-Nitrobenzylidenyl)-5-iodo-2-indolinone **204001-63-8P**,  
 3-(3,5-Dibromo-2-hydroxybenzylidenyl)-5-iodo-2-indolinone **204001-64-9P**, 3-(3-Fluoro-2-hydroxybenzylidenyl)-5-iodo-2-indolinone **204001-65-0P**, 3-(3-Bromo-4-hydroxybenzylidenyl)-5-iodo-2-indolinone **204001-66-1P**, 3-(4-tert-Butylbenzylidenyl)-5-iodo-2-indolinone **204001-67-2P**, 3-[(2-Bromothien-5-yl)methylidenyl]-5-iodo-2-indolinone **204001-68-3P**,  
 3-(3-Ethoxy-2-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone **204001-69-4P**, 3-(3,5-Dichloro-2-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone **204001-70-7P**, 3-(5-Chloro-2-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone **204001-71-8P**,  
 3-[4-(Diethylamino)-2-hydroxybenzylidenyl]-5-bromo-4-methyl-2-indolinone **204001-72-9P**, 3-(4-Nitrobenzylidenyl)-5-bromo-4-methyl-2-indolinone **204001-73-0P**, 3-(3,5-Dibromo-2-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone **204001-74-1P**, 3-(3-Fluoro-2-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone **204001-75-2P**,  
 3-(3-Bromo-4-hydroxybenzylidenyl)-5-bromo-4-methyl-2-indolinone **204001-76-3P**, 3-(4-tert-Butylbenzylidenyl)-5-bromo-4-methyl-2-indolinone **204001-77-4P**, 3-[(2-Bromothien-5-yl)methylidenyl]-5-bromo-4-methyl-2-indolinone **204001-78-5P**,  
 3-(3-Ethoxy-2-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone **204001-79-6P**, 3-(3,5-Dichloro-2-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone **204001-80-9P**,  
 3-(5-Chloro-2-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone **204001-81-0P**, 3-[4-(Diethylamino)-2-hydroxybenzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone **204001-82-1P**,  
 3-(4-Nitrobenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone **204001-83-2P**, 3-(3,5-Dibromo-2-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone **204001-84-3P**,  
 3-(3-Fluoro-2-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone **204001-85-4P**, 3-(3-Bromo-4-hydroxybenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone **204001-86-5P**,  
 3-(4-tert-Butylbenzylidenyl)-5-[(methylamino)sulfonyl]-2-indolinone **204001-87-6P**, 3-[(2-Bromothien-5-yl)methylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone **204001-88-7P**,  
 3-(3-Ethoxy-2-hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204001-89-8P**, 3-(3,5-Dichloro-2-hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204001-90-1P**,  
 3-(5-Chloro-2-hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204001-91-2P**, 3-[4-(Diethylamino)-2-hydroxybenzylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204001-92-3P**,  
 3-(4-Nitrobenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204001-93-4P**, 3-(3,5-Dibromo-2-hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204001-94-5P**,  
 3-(3-Fluoro-2-hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204001-95-6P**, 3-(3-Bromo-4-hydroxybenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204001-96-7P**,  
 3-(4-tert-Butylbenzylidenyl)-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204001-97-8P**, 3-[(2-Bromothien-5-yl)methylidenyl]-5-[[[4-(trifluoromethyl)phenyl]amino]sulfonyl]-2-indolinone **204001-98-9P**,  
 3-(3-Ethoxy-2-hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **204001-99-0P**, 3-(3,5-Dichloro-2-hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **204002-00-6P**,  
 3-(5-Chloro-2-hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone **204002-01-7P**, 3-[4-(Diethylamino)-2-hydroxybenzylidenyl]-5-

(morpholinosulfonyl)-2-indolinone 204002-02-8P,  
 3-(4-Nitrobenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone  
 204002-03-9P, 3-(3,5-Dibromo-2-hydroxybenzylidenyl)-5-  
 (morpholinosulfonyl)-2-indolinone 204002-04-0P,  
 3-(3-Fluoro-2-hydroxybenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone  
 204002-05-1P, 3-(3-Bromo-4-hydroxybenzylidenyl)-5-  
 (morpholinosulfonyl)-2-indolinone 204002-06-2P,  
 3-(4-tert-Butylbenzylidenyl)-5-(morpholinosulfonyl)-2-indolinone  
 204002-07-3P, 3-[(2-Bromothien-5-yl)methylidenyl]-5-  
 (morpholinosulfonyl)-2-indolinone 204002-08-4P,  
 3-(3-Ethoxy-2-hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone  
 204002-09-5P, 3-(3,5-Dichloro-2-hydroxybenzylidenyl)-5-(2-  
 chloroethyl)-2-indolinone 204002-10-8P, 3-(5-Chloro-2-  
 hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 204002-11-9P  
 , 3-[4-(Diethylamino)-2-hydroxybenzylidenyl]-5-(2-chloroethyl)-2-  
 indolinone 204002-12-0P, 3-(4-Nitrobenzylidenyl)-5-(2-  
 chloroethyl)-2-indolinone 204002-13-1P, 3-(3,5-Dibromo-2-  
 hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone 204002-14-2P  
 , 3-(3-Fluoro-2-hydroxybenzylidenyl)-5-(2-chloroethyl)-2-indolinone  
 204002-15-3P, 3-(3-Bromo-4-hydroxybenzylidenyl)-5-(2-chloroethyl)-  
 2-indolinone 204002-16-4P, 3-(4-tert-Butylbenzylidenyl)-5-(2-  
 chloroethyl)-2-indolinone 204002-17-5P, 3-[(2-Bromothien-5-  
 yl)methylidenyl]-5-(2-chloroethyl)-2-indolinone 204003-31-6P  
 204003-32-7P 204003-42-9P 204003-84-9P  
 204003-85-0P 204003-88-3P 204003-89-4P  
 204003-90-7P 204003-91-8P 204003-96-3P  
 204003-97-4P 204004-10-4P 204004-11-5P  
 204004-29-5P 204004-86-4P 204004-92-2P  
 204004-94-4P 204005-03-8P 204005-21-0P  
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 204005-54-9P 204005-56-1P 204005-58-3P  
 204005-59-4P 204005-60-7P 204005-66-3P  
 204005-67-4P 204005-71-0P 204005-75-4P  
 204005-76-5P 204005-90-3P 204006-01-9P  
 204006-02-0P 204006-03-1P 204006-04-2P  
 204006-05-3P 204006-06-4P 204006-19-9P  
 204006-26-8P 204006-35-9P 204006-83-7P  
 204006-93-9P 204006-96-2P 204006-98-4P  
 204007-13-6P, 3-(4-Hydroxy-3-methylbenzylidenyl)-5-bromo-4-methyl-  
 2-indolinone

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic  
 preparation); THU (Therapeutic use); BIOL (Biological study); PREP  
 (Preparation); USES (Uses)

(prepn. and testing of indolinone combinatorial library as protein  
 kinase inhibitors)

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ACCESSION NUMBER: 1998:1471 HCAPLUS

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TITLE: Preparation of substituted quinolylmethylenoxindole  
 analogs as tyrosine kinase inhibitors

INVENTOR(S): Battistini, Carlo; Ermoli, Antonella; Vioglio, Sergio;  
 Buzzetti, Franco; Ballinari, Dario

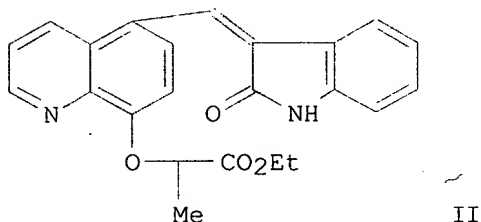
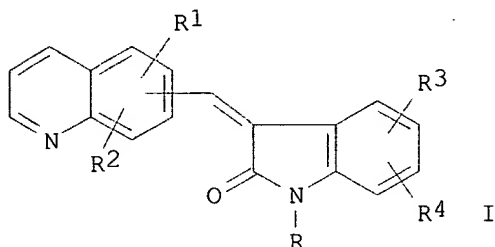
PATENT ASSIGNEE(S): Pharmacia + Upjohn S.p.A., Italy; Battistini, Carlo;  
 Ermoli, Antonella; Vioglio, Sergio; Buzzetti, Franco;  
 Ballinari, Dario

SOURCE: PCT Int. Appl., 51 pp.  
 CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9746551	A1	19971211	WO 1997-EP2673	19970515
W: JP, US				
RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
EP 876365	A1	19981111	EP 1997-927035	19970515
R: DE, GB, IT				
JP 11510823	T2	19990921	JP 1997-500166	19970515
US 5905149	A	19990518	US 1998-983516	19980129
PRIORITY APPLN. INFO.:			GB 1996-11797	19960606
			WO 1997-EP2673	19970515
OTHER SOURCE(S):		MARPAT 128:61437		
GI				



AB The title compds. [I; R1-R4 = X(CH<sub>2</sub>)<sub>m</sub>NH<sub>2</sub>, X(CH<sub>2</sub>)<sub>m</sub>NR<sub>5</sub>R<sub>6</sub>, etc.; R = H, (CH<sub>2</sub>)<sub>n</sub>COR<sub>7</sub>, etc.; n = 1-4; m = 2-4; R<sub>5</sub>, R<sub>6</sub> = H, C1-6 alkyl; R<sub>7</sub> = (un)substituted aminoacids, etc.] and the pharmaceutically acceptable salts thereof are prepd. I, possessing tyrosine kinase inhibitory activity, are useful as immunomodulating agents, and antimetastatic and anticancer agents, or in the control of **angiogenesis** and atheromatous plaque, and treatment of Alzheimer's disease. Thus, 8-hydroxyquinoline-5-carbaldehyde was reacted with 2-oxoindole in the presence of piperidine and then reacted with MeCHBrCO<sub>2</sub>Et in the presence of Bu<sub>4</sub>NF to give the title compd. (II), which showed IC<sub>50</sub> of 39.5 .mu.M against K562 cell growth in vivo. A formulation contg. I were also prepd.

IT 200285-07-0P 200285-08-1P 200285-09-2P  
 200285-10-5P 200285-11-6P 200285-12-7P  
 200285-13-8P 200285-14-9P 200285-15-0P  
 200285-20-7P 200285-21-8P 200285-22-9P  
 RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP

(Preparation); USES (Uses)

(prepn. of substituted quinolylmethylenoxindole analogs as tyrosine kinase inhibitors)

IT 137479-19-7P 168464-10-6P 200285-25-2P  
200285-26-3PRL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation)  
(prepn. of substituted quinolylmethylenoxindole analogs as tyrosine kinase inhibitors)

L18 ANSWER 15 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1997:805721 HCAPLUS

DOCUMENT NUMBER: 128:61424

TITLE: Preparation of substituted tetralinylmethylen-2-oxindole analogs as tyrosine kinase inhibitors

INVENTOR(S): Battistini, Carlo; Ermoli, Antonella; Vioglio, Sergio; Buzzetti, Franco; Ballinari, Dario

PATENT ASSIGNEE(S): Pharmacia &amp; Upjohn, S.p.A., Italy; Battistini, Carlo; Ermoli, Antonella; Vioglio, Sergio; Buzzetti, Franco; Ballinari, Dario

SOURCE: PCT Int. Appl., 43 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

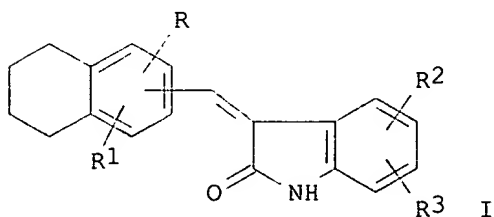
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9745409	A1	19971204	WO 1997-EP2672	19970515
W: JP, US				
RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
EP 853614	A1	19980722	EP 1997-927034	19970515
EP 853614	B1	20011004		
R: DE, GB, IT				
JP 11510822	T2	19990921	JP 1997-541580	19970515
US 6147073	A	20001114	US 1998-981473	19980112
PRIORITY APPLN. INFO.:			GB 1996-10964	A 19960524
			WO 1997-EP2672	W 19970515

OTHER SOURCE(S): MARPAT 128:61424

GI



AB The title compds. [I; R, R1-R3 = X(CH<sub>2</sub>)<sub>m</sub>NH<sub>2</sub>, X(CH<sub>2</sub>)<sub>m</sub>NR<sub>4</sub>R<sub>5</sub>, etc.; X = O, S, NH, etc.; m = 2-4; R<sub>4</sub>, R<sub>5</sub> = H, C1-6 alkyl, etc.] and pharmaceutically acceptable salts thereof are prepd. I, possessing tyrosine kinase inhibitory activity, are useful as antiproliferative, anti-metastatic, immunomodulating, and anticancer agents, or in the control of **angiogenesis** and in the treatment of Alzheimer's diseases. Thus,

I (R = R1 = R3 = H, R2 = 5-NH2) (prepn. given) was reacted with N-tert-butoxycarbonyl-L-glutamic acid tert-Bu ester in the presence of benzotriazol-1-yloxytripyrrolidinophosphonium hexafluorophosphate and N-methylmorpholine, and then treated with CF3CO2H to give 40% I.CF3CO2H (R, R1, R3 = same as above, R2 = glutamylamino), which showed IC50 of 5.97 .mu.M against K562 cell growth in vivo. A formulation contg. I were prepd.

IT 200191-17-9P 200191-19-1P 200191-21-5P  
200191-23-7P 200191-25-9P 200191-27-1P  
200191-29-3P

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of substituted tetralinylmethylen-2-oxoindole analogs as tyrosine kinase **inhibitors**)

IT 200191-30-6P 200191-34-0P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation)  
(prepn. of substituted tetralinylmethylen-2-oxoindole analogs as tyrosine kinase **inhibitors**)

L18 ANSWER 16 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1996:746204 HCAPLUS

DOCUMENT NUMBER: 126:18783

TITLE: Substituted indolylmethylen-oxindole analogs as tyrosine kinase inhibitors

INVENTOR(S): Battistini, Carlo; Ballinari, Dario; Ermoli, Antonella; Penco, Sergio; Vioglio, Sergio

PATENT ASSIGNEE(S): Pharmacia S.P.A., Italy

SOURCE: PCT Int. Appl., 53 pp.

CODEN: PIXXD2

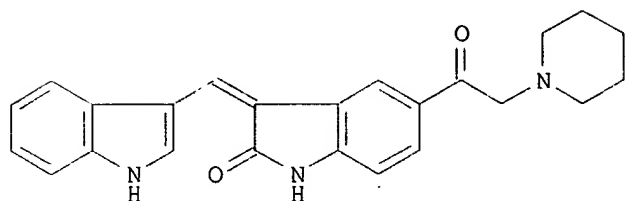
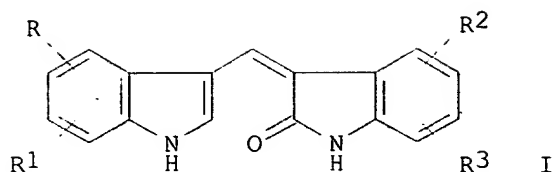
DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9632380	A1	19961017	WO 1996-EP1165	19960314
W: JP, US				
RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
EP 764152	A1	19970326	EP 1996-907500	19960314
R: DE, ES, FR, GB, IT, SE				
JP 10501821	T2	19980217	JP 1996-530667	19960314
US 5849710	A	19981215	US 1996-750208	19961204
PRIORITY APPLN. INFO.:			GB 1995-7298	19950407
			WO 1996-EP1165	19960314
OTHER SOURCE(S):			MARPAT 126:18783	
GI				



II

- AB Indol-3-ylmethylene-2-oxindole derivs. I and their pharmaceutically acceptable salts are disclosed [wherein 1 or 2 of R, R1, R2, and R3 = X(CH2)mNH2, X(CH2)mNR4R5, X(CH2)mNHR6, NHC(:NH)NH2, NHC(:NH)NR4R5, NHC(:NH)NHR6, N:CHNH2, N:CHNR4R5, N:CHNHR6, X(CH2)mCOR7, CORa, COR8, YCOY'R9, NHR6, NHR10 group; remaining groups within R and R1-R3 = H, halo, amino, OH, alkyl, alkoxy, CO2H, alkoxycarbonyl, alkanoyloxy, cyano, NR4R5; X = O, S, NH; m = 1-4; 1 of R4 and R5 = H or alkyl, and other = alkyl; or NR4R5 forms satd. monoheterocycle; R6 = alkanoyl, 1- to 3-residue (un)substituted peptidyl; R7 = OH, amino, alkoxy, NR4R5; Ra = amino terminus of 1- to 3-unit peptidyl; R8 = alkoxy, phenylalkoxy, (CH2)nNH2, (CH2)nNR4R5, (CH2)nNHR6; n = 1-2; Y, Y' = NH, O; R9 = Ph, alkyl, phenylalkyl; R10 = mono-, di- or trihydroxyalkyl]. I have tyrosine kinase inhibiting activity, and are useful as antiproliferative, antimetastatic, anticancer, antiatheromatous, anti-Alzheimer, and immunomodulating agents. For example, 2-indolinone reacted with BrCH2COBr and AlCl3 to give the 5-(2-bromoacetyl) deriv., which underwent amination with piperidine and then condensation with indole-3-carboxaldehyde, to give title compd. II (FCE 28484). In tests for inhibition of p45 v-abl kinase and K562 leukemia cells in vitro, II had IC50 of 0.78 and 4.82 .mu.M, resp.
- IT 168464-17-3P 184021-39-4P 184021-56-5P  
184021-79-2P 184021-85-0P 184021-97-4P  
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation)  
(intermediate; prepn. of (indolylmethylene)oxindole analogs as tyrosine kinase inhibitors)
- IT 184020-98-2P  
RL: BAC (Biological activity or effector, except adverse); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
(prepn. of (indolylmethylene)oxindole analogs as tyrosine kinase inhibitors)
- IT 181223-99-4P 184020-79-9P 184020-86-8P  
184020-93-7P 184021-06-5P 184021-15-6P  
184021-23-6P  
RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
(prepn. of (indolylmethylene)oxindole analogs as tyrosine kinase inhibitors)
- IT 184020-69-7P  
RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP

(Preparation); USES (Uses)  
 (prepn. of (indolylmethylene)oxindoles as tyrosine kinase  
 inhibitors)

L18 ANSWER 17 OF 17 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1995:828284 HCAPLUS

DOCUMENT NUMBER: 123:227985

TITLE: Arylidene and heteroarylidene oxindole derivatives as  
 tyrosine kinase inhibitors

INVENTOR(S): Buzzetti, Franco; Longo, Antonio; Brasca, Maria  
 Gabriella; Orzi, Fabrizio; Crugnola, Angelo;  
 Ballinari, Dario; Mariani, Mariangela

PATENT ASSIGNEE(S): Farmitalia Carlo Erba S.r.l., Italy

SOURCE: PCT Int. Appl., 62 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

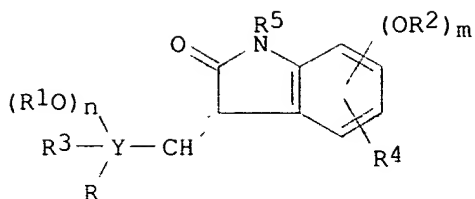
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9501349	A1	19950112	WO 1994-EP1715	19940526
W: AU, BB, BG, BR, BY, CA, CN, CZ, FI, HU, JP, KP, KR, KZ, LV, MG, MN, MW, NO, NZ, PL, RO, RU, SD, SK, UA, UZ, VN				
RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
CA 2142472	AA	19950112	CA 1994-2142472	19940526
AU 9469719	A1	19950124	AU 1994-69719	19940526
AU 679754	B2	19970710		
EP 658159	A1	19950621	EP 1994-918379	19940526
EP 658159	B1	20000823		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, NL, PT, SE				
CN 1111454	A	19951108	CN 1994-190452	19940526
JP 08500847	T2	19960130	JP 1994-503150	19940526
HU 72047	A2	19960328	HU 1995-954	19940526
EP 987263	A2	20000322	EP 1999-203366	19940526
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, NL, SE, PT, IE				
AT 195734	E	20000915	AT 1994-918379	19940526
ES 2152317	T3	20010201	ES 1994-918379	19940526
US 5656654	A	19970812	US 1994-263666	19940622
ZA 9404730	A	19950713	ZA 1994-4730	19940630
FI 9500859	A	19950224	FI 1995-859	19950224
PRIORITY APPLN. INFO.:			GB 1993-13638	A 19930701
			EP 1994-918379	A3 19940526
			WO 1994-EP1715	W 19940526

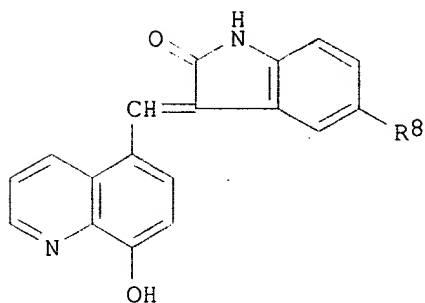
OTHER SOURCE(S): MARPAT 123:227985

GI





I



II

AB Title derivs. I [Y = naphthalene, tetralin, quinoline or isoquinoline system; R = H, plus oxo when Y is tetralin;  $R^1$ ,  $R^2$  independently = H, C1-6 alkyl or C2-6 alkanoyl;  $m = 0-2$ ;  $n = 0-3$ ;  $R^3$  independently = H, halo, cyano, C1-6 alkyl, carboxy, nitro or  $NR^6R^7$  where  $R^6$ ,  $R^7$  independently = H, C1-6 alkyl;  $R^5$  = H, C1-6 alkyl] and their pharmaceutically acceptable salts, which are useful as tyrosine kinase inhibitors, are claimed. The E- and Z-isomers of approx. 85 compds. are specifically claimed. Several synthetic examples are given. For example, condensation of 8-hydroxyquinoline-5-carboxaldehyde with 5-hydroxy-2-oxindole in EtOH in the presence of piperidine at 60-70.degree. gave 60% title compd. II ( $R^8$  = OH). Among test results for 10 selected I for inhibition of p45 v-abl kinase in vitro, and for inhibition of cultured K562 human leukemia cell growth, II ( $R^8$  = Br) had  $IC_{50}$  values of 2.6 and 0.62  $\mu M$ , resp.

IT 149492-63-7P 168462-84-8P 168462-85-9P  
 168462-86-0P 168462-87-1P 168462-88-2P  
 168462-89-3P 168462-90-6P 168462-91-7P  
 168462-92-8P 168462-93-9P 168462-94-0P  
 168462-95-1P 168462-96-2P 168462-97-3P  
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 168464-22-0P 168464-23-1P

RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(invention compd.; prepn. of (hetero)arylidene oxindole derivs. as tyrosine kinase inhibitors)

IT 168464-07-1 168464-08-2 168464-09-3

RL: RCT (Reactant)

(starting material; prepn. of (hetero)arylidene oxindole derivs. as tyrosine kinase inhibitors)

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 E82 THROUGH E999 ASSIGNED

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STRUCTURE FILE UPDATES: 14 OCT 2001 HIGHEST RN 362009-74-3  
 DICTIONARY FILE UPDATES: 14 OCT 2001 HIGHEST RN 362009-74-3

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 HELP CROSSOVER for details.

Calculated physical property data is now available. See HELP PROPERTIES  
 for more information. See STNote 27, Searching Properties in the CAS  
 Registry File, for complete details:  
<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

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E# OR SYSTEM LIMIT REACHED WHILE PROCESSING ANSWER 13  
E1 THROUGH E999 ASSIGNED

=> del select y

=> select hit rn l18 13

E# OR SYSTEM LIMIT REACHED WHILE PROCESSING ANSWER 13  
E1 THROUGH E999 ASSIGNED

=> delete select y

ALL E# DEFINITIONS DELETED

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E1 THROUGH E163 ASSIGNED

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DICTIONARY FILE UPDATES: 14 OCT 2001 HIGHEST RN 362009-74-3

TSCA INFORMATION NOW CURRENT THROUGH July 7, 2001

Please note that search-term pricing does apply when  
conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER see  
HELP CROSSOVER for details.

Calculated physical property data is now available. See HELP PROPERTIES  
for more information. See STNote 27, Searching Properties in the CAS  
Registry File, for complete details:  
<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

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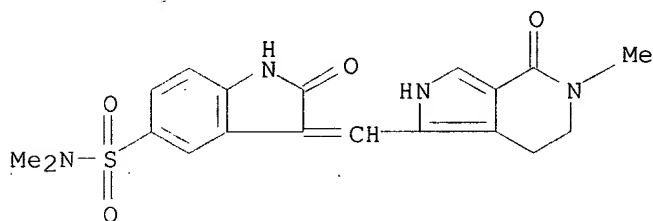
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L20 1080 S E1-E163 OR L19  
L21 1079 S L5 AND L20

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=>

=> d ide can 121 1 100 200 300 400 500 600 700 800 900 1000 1079

L21 ANSWER 1 OF 1079 REGISTRY COPYRIGHT 2001 ACS  
RN 358733-27-4 REGISTRY  
CN 1H-Indole-5-sulfonamide, 2,3-dihydro-N,N-dimethyl-2-oxo-3-[(4,5,6,7-tetrahydro-5-methyl-4-oxo-2H-pyrrolo[3,4-c]pyridin-1-yl)methylene]- (9CI)  
(CA INDEX NAME)  
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MF C19 H20 N4 O4 S  
SR CA  
LC STN Files: CA, CAPLUS



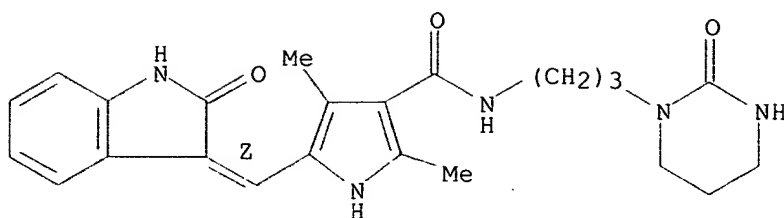
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1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 135:226880

L21 ANSWER 100 OF 1079 REGISTRY COPYRIGHT 2001 ACS  
RN 356069-30-2 REGISTRY  
CN 1H-Pyrrole-3-carboxamide, 5-[(Z)-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2,4-dimethyl-N-[3-(tetrahydro-2-oxo-1(2H)-pyrimidinyl)propyl]- (9CI) (CA INDEX NAME)  
FS STEREOSEARCH  
MF C23 H27 N5 O3  
SR CA  
LC STN Files: CA, CAPLUS

Double bond geometry as shown.

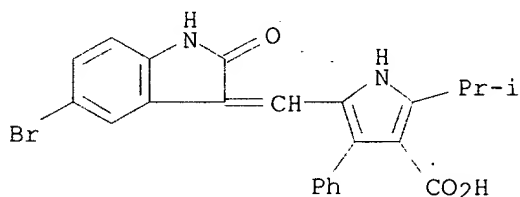


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1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 135:195497

L21 ANSWER 200 OF 1079 REGISTRY COPYRIGHT 2001 ACS  
RN **342641-53-6** REGISTRY  
CN 1H-Pyrrole-3-carboxylic acid, 5-[(5-bromo-1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-2-(1-methylethyl)-4-phenyl- (9CI) (CA INDEX NAME)  
FS 3D CONCORD  
MF C23 H19 Br N2 O3  
SR CA  
LC STN Files: CA, CAPLUS



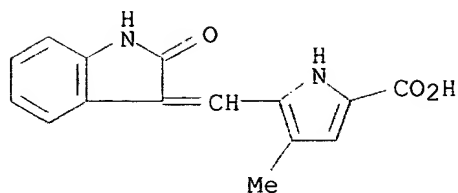
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

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2 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 135:195497

REFERENCE 2: 135:19549

L21 ANSWER 300 OF 1079 REGISTRY COPYRIGHT 2001 ACS  
RN **280748-39-2** REGISTRY  
CN 1H-Pyrrole-2-carboxylic acid, 5-[(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-4-methyl- (9CI) (CA INDEX NAME)  
OTHER NAMES:  
CN SU 6595  
FS 3D CONCORD  
MF C15 H12 N2 O3  
SR CA  
LC STN Files: CA, CAPLUS, TOXLIT



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

5 REFERENCES IN FILE CA (1967 TO DATE)  
1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA  
5 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 135:195497

REFERENCE 2: 135:107247

REFERENCE 3: 135:19549

REFERENCE 4: 134:141334

REFERENCE 5: 133:84238

L21 ANSWER 400 OF 1079 REGISTRY COPYRIGHT 2001 ACS

RN 210303-50-7 REGISTRY

CN 1H-Pyrrole-3-propanoic acid, 5-[(Z)-(1,2-dihydro-2-oxo-3H-indol-3-ylidene)methyl]-4-methyl- (9CI) (CA INDEX NAME)

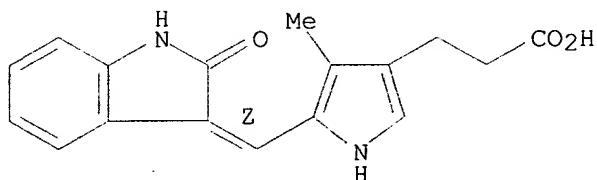
FS STEREOSEARCH

MF C17 H16 N2 O3

SR CA

LC STN Files: CA, CAPLUS, TOXLIT

Double bond geometry as shown.



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1967 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 129:117426

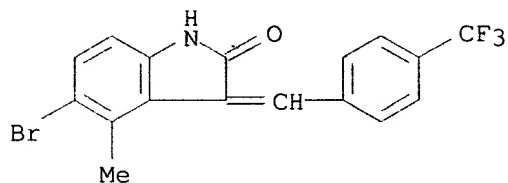
L21 ANSWER 500 OF 1079 REGISTRY COPYRIGHT 2001 ACS

RN 203992-55-6 REGISTRY

CN 2H-Indol-2-one, 5-bromo-1,3-dihydro-4-methyl-3-[[4-(trifluoromethyl)phenyl]methylene]- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 3-[4-(Trifluoromethyl)benzylidene]-5-bromo-4-methyl-2-indolinone  
 MF C17 H11 Br F3 N O  
 SR CA  
 LC STN Files: CA, CAPLUS, USPATFULL

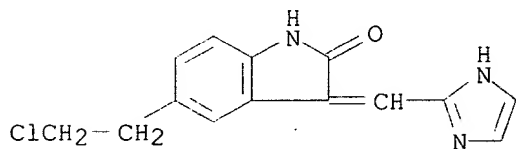


\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1967 TO DATE)  
 1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 128:204803

L21 ANSWER 600 OF 1079 REGISTRY COPYRIGHT 2001 ACS  
 RN 203991-55-3 REGISTRY  
 CN 2H-Indol-2-one, 5-(2-chloroethyl)-1,3-dihydro-3-(1H-imidazol-2-ylmethylene)- (9CI) (CA INDEX NAME)  
 OTHER NAMES:  
 CN 3-[(Imidazol-2-yl)methylene]-5-(2-chloroethyl)-2-indolinone  
 FS 3D CONCORD  
 MF C14 H12 Cl N3 O  
 SR CA  
 LC STN Files: CA, CAPLUS, USPATFULL



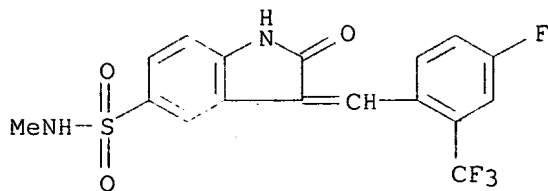
\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1967 TO DATE)  
 1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 128:204803

L21 ANSWER 700 OF 1079 REGISTRY COPYRIGHT 2001 ACS  
 RN 203990-24-3 REGISTRY  
 CN 1H-Indole-5-sulfonamide, 3-[[4-fluoro-2-(trifluoromethyl)phenyl]methylene]-2,3-dihydro-N-methyl-2-oxo- (9CI) (CA INDEX NAME)  
 OTHER NAMES:  
 CN 3-[4-Fluoro-2-(trifluoromethyl)benzylidenyl]-5-[(methylamino)sulfonyl]-2-indolinone

FS 3D CONCORD  
 MF C17 H12 F4 N2 O3 S  
 SR CA  
 LC STN Files: CA, CAPLUS, USPATFULL

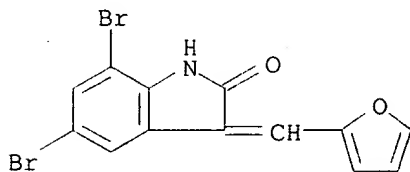


\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1967 TO DATE)  
 1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 128:204803

L21 ANSWER 800 OF 1079 REGISTRY COPYRIGHT 2001 ACS  
 RN 203989-13-3 REGISTRY  
 CN 2H-Indol-2-one, 5,7-dibromo-3-(2-furanylmethylene)-1,3-dihydro- (9CI) (CA INDEX NAME)  
 OTHER NAMES:  
 CN 3-(Furan-2-ylmethylydenyl)-5,7-dibromo-2-indolinone  
 FS 3D CONCORD  
 MF C13 H7 Br2 N O2  
 SR CA  
 LC STN Files: CA, CAPLUS, USPATFULL



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1967 TO DATE)  
 1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 128:204803

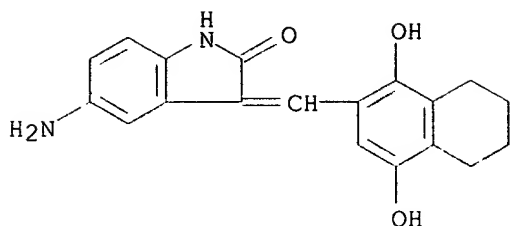
L21 ANSWER 900 OF 1079 REGISTRY COPYRIGHT 2001 ACS  
 RN 200191-19-1 REGISTRY  
 CN 2H-Indol-2-one, 5-amino-1,3-dihydro-3-[(5,6,7,8-tetrahydro-1,4-dihydroxy-2-naphthalenyl)methylene]-, mono(trifluoroacetate) (salt) (9CI) (CA INDEX NAME)  
 MF C19 H18 N2 O3 . C2 H F3 O2  
 SR CA  
 LC STN Files: CA, CAPLUS, USPATFULL



CM 1

CRN 200191-18-0

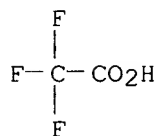
CMF C19 H18 N2 O3



CM 2

CRN 76-05-1

CMF C2 H F3 O2



1 REFERENCES IN FILE CA (1967 TO DATE)

1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 128:61424

L21 ANSWER 1000 OF 1079 REGISTRY COPYRIGHT 2001 ACS

RN 168463-44-3 REGISTRY

CN 2H-Indol-2-one, 1,3-dihydro-5-hydroxy-3-[(5-methoxy-1H-indol-3-yl)methylene]-, (E)- (9CI) (CA INDEX NAME)

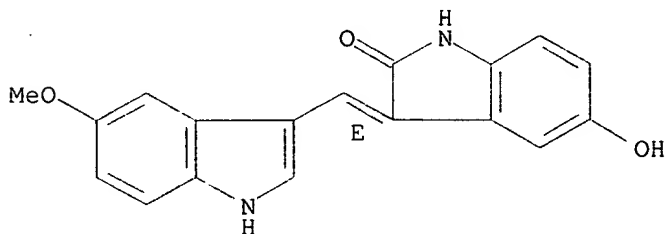
FS STEREOSEARCH

MF C18 H14 N2 O3

SR CA

LC STN Files: CA, CAPLUS, USPATFULL

Double bond geometry as shown.

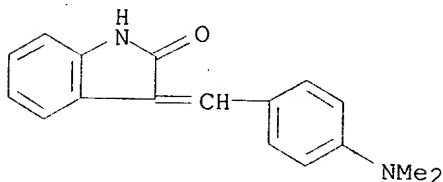


\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1 REFERENCES IN FILE CA (1967 TO DATE)  
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 123:227985

L21 ANSWER 1079 OF 1079 REGISTRY COPYRIGHT 2001 ACS  
RN 5812-07-7 REGISTRY  
CN 2H-Indol-2-one, 3-[[4-(dimethylamino)phenyl]methylene]-1,3-dihydro- (9CI)  
(CA INDEX NAME)  
OTHER CA INDEX NAMES:  
CN 2-Indolinone, 3-[p-(dimethylamino)benzylidene]- (7CI, 8CI)  
OTHER NAMES:  
CN 3-(4-Dimethylaminobenzylidenyl)-2-indolinone  
CN SU 4312  
MF C17 H16 N2 O  
LC STN Files: BEILSTEIN\*, BIOSIS, CA, CAOLD, CAPLUS, CHEMCATS, TOXLIT,  
USPATFULL  
(\*File contains numerically searchable property data)



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

13 REFERENCES IN FILE CA (1967 TO DATE)  
13 REFERENCES IN FILE CAPLUS (1967 TO DATE)  
1 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 133:234344  
REFERENCE 2: 130:348928  
REFERENCE 3: 130:252240  
REFERENCE 4: 130:237471  
REFERENCE 5: 130:237470  
REFERENCE 6: 130:149064  
REFERENCE 7: 129:330650  
REFERENCE 8: 129:175549  
REFERENCE 9: 127:314804  
REFERENCE 10: 126:139901